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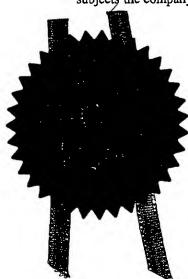
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Dated 22 February 2001

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Description 27

Claim(s) 5

Abstract

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Statement of inventorship and right to grant of a patent (Patents Form 7/77)

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DRUG TARGETS IN CANDIDA ALBICANS

The present invention is concerned with the identification of genes or functional fragments thereof from Candida albicans which are critical for growth and cell division and which genes may be used as selective drug targets to treat Candida albicans associated infections. Novel nucleic acid sequences from Candida albicans are also provided and which encode the polypeptides which are critical for growth of Candida albicans.

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Opportunistic infections in immunocompromised hosts represent an increasingly common cause of mortality and morbidity. Candida species are among the most commonly identified fungal pathogens associated with such opportunistic infections, with Candida albicans being the most common species. Such fungal infections are thus problematical in, for example, AIDS populations in addition to normal healthy women where Candida albicans yeasts represent the most common cause of vulvovaginitis.

Although compounds do exist for treating such disorders, such as for example, amphotericin, these drugs are generally limited in their treatment because of their toxicity and side effects. Therefore, there exists a need for new compounds which may be used to treat Candida associated infections in addition to compounds which are selective in their action against Candida albicans.

Classical approaches for identifying anti-fungal compounds have relied almost exclusively on inhibition of fungal or yeast growth as an endpoint. Libraries of natural products, semi-synthetic, or synthetic chemicals are screened for their ability to kill or arrest growth of the target pathogen or a related

nonpathogenic model organism. These tests are cumbersome and provide no information about a compounds mechanism of action. The promising lead compounds that emerge from such screens must then be tested for possible host-toxicity and detailed mechanism of action studies must subsequently be conducted to identify the affected molecular target.

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The present inventors have now identified a range of nucleic acid sequences from Candida albicans which encode polypeptides which are critical for its survival and growth. These sequences represent novel targets which can be incorporated into an assay to selectively identify compounds capable of inhibiting expression of such polypeptides and their potential use in alleviating diseases or conditions associated with Candida albicans infection.

Therefore, according to a first aspect of the invention there is provided a nucleic acid molecule encoding a polypeptide which is critical for survival and growth of the yeast Candida albicans and which nucleic acid molecule comprises any of the sequences of nucleotides illustrated in Figures 1, 2, 4 to 7, 9 to 11, 13, 15 to 20, 22 to 26, 28 to 32, 34 to 43, 45a and b, 47 to 49, 51, 52, 53 to 57, 59 and 60.

A further aspect of the invention comprises a nucleic acid molecule encoding a polypeptide which is critical for survival and growth of the yeast Candida albicans and which nucleic acid molecule comprises any of the sequences of nucleotides illustrated in Figure 1, 2, 36, 37a and b, 38, 39 and 40 and fragments or derivatives of said nucleic acid molecules.

Letters utilised in the sequences according to the invention which are not recognisable as letters of the genetic code signify a position in the nucleic acid sequence where one or more of bases A, G, C or T can occupy the nucleotide position. Representative letters used to identify the range of bases which can be used are as follows:

5	M:	A or C
	R:	A or G
	W:	A or T
	s:	C or G
	Y:	C or T
10	K:	G or T
	v:	A or C or G
	H:	A or C or T
	D:	A or G or T
	B :	C or G or T
15	N:	G or A or T or C

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In one embodiment of each of the above identified aspects of the invention the nucleic acid may comprise a mRNA molecule or alternatively a DNA and preferably a cDNA molecule.

Also provided by the present invention is a nucleic acid molecule capable of hybridising to the nucleic acid molecules illustrated in any of Figures 1 to 61 under high stringency conditions.

Stringency of hybridisation as used herein refers to conditions under which polynucleic acids are stable. The stability of hybrids is reflected in the melting temperature (Tm) of the hybrids. Tm can be approximated by the formula:

81.5°C+16.6(log₁₀[Na⁺]+0.41 (%G&C)-6001/1

wherein 1 is the length of the hybrids in nucleotides. Tm decreases approximately by 1-1.5°C with every 1% decrease in sequence homology.

The nucleic acid capable of hybridising to nucleic acid molecules according to the invention will generally be at least 70%, preferably at least 80 or 90% and more preferably at least 95% homologous to the nucleotide sequences illustrated in any of Figures 1 to 61.

The DNA molecules according to the invention may, advantageously, be included in a suitable expression vector to express polypeptides encoded therefrom in a suitable host.

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An expression vector according to the invention includes a vector having a nucleic acid according to the invention operably linked to regulatory sequences, such as promoter regions, that are capable of effecting expression of said DNA fragments. The term "operably linked" refers to a juxta position wherein the components described are in a relationship permitting them to function in their intended manner. Such vectors may be transformed into a suitable host cell to provide for expression of a polypeptide according to the invention. Thus, in a further aspect, the invention provides a process for preparing polypeptides according to the invention which comprises cultivating a host cell, transformed or transfected with an expression vector as described above under conditions to provide for expression by the vector of a coding sequence encoding the polypeptides, and recovering the expressed polypeptides.

The vectors may be, for example, plasmid, virus or phage vectors provided with an origin of replication, optionally a promoter for the expression of said nucleotide and optionally a regulator of the promoter. The vectors may contain one or more selectable markers, such as, for example, ampicillin

resistance.

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Polynucleotides according to the invention may be inserted into the vectors described in an antisense orientation in order to provide for the production of antisense RNA. Antisense RNA or other antisense nucleic acids may be produced by synthetic means.

In accordance with the present invention, a defined nucleic acid includes not only the identical nucleic acid but also any minor base variations including in particular, substitutions in bases which result in a synonymous codon (a different codon specifying the same amino acid residue) due to the degenerate code in conservative amino acid substitutions. The term "nucleic acid sequence" also includes the complementary sequence to any single stranded sequence given regarding base variations.

The present invention also comprises within its scope proteins or polypeptides expressed by the nucleic acid molecules according to the invention or a functional equivalent, derivative or bioprecursor thereof.

The present invention also advantageously provides nucleic acid sequences of at least approximately 15 contiguous nucleotides of a nucleic acid according to the invention and preferably from 15 to 50 nucleotides. These sequences may, advantageously be used as probes or primers to initiate replication, or the like. Such nucleic acid sequences may be produced according to techniques well known in the art, such as by recombinant or synthetic means. They may also be used in diagnostic kits or the like for detecting the presence of a nucleic acid according to the invention. These tests generally comprise contacting the probe with the sample under hybridising conditions and detecting for the presence

of any duplex or triplex formation between the probe and any nucleic acid in the sample.

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Advantageously, the nucleic acid sequences, according to the invention may be produced using such recombinant or synthetic means, such as for example using PCR cloning mechanisms which generally involve making a pair of primers, which may be from approximately 15 to 50 nucleotides to a region of the gene which is desired to be cloned, bringing the primers into contact with mRNA, cDNA, or genomic DNA from a human cell, performing a polymerase chain reaction under conditions which bring about amplification of the desired region, isolated the amplified region or fragment and recovering the amplified DNA. Generally, such techniques as defined herein are well known in the art, such as described in Sambrook et al (Molecular Cloning: a Laboratory Manual, 1989).

The nucleic acids or oligonucleotides according to the invention may carry a revealing label. Suitable labels include radioisotopes such as ³²P or ³⁹S, enzyme labels or other protein labels such as biotin or fluorescent markers. Such labels may be added to the nucleic acids or oligonucleotides of the invention and may be detected using known techniques per se.

The polypeptide or protein according to the invention includes all possible amino acid variants encoded by the nucleic acid molecule according to the invention including a polypeptide encoded by said molecule and having conservative amino acid changes. Polypeptides according to the invention further include variants of such sequences, including naturally occurring allelic variants which are substantially homologous to said polypeptides. In

this context, substantial homology is regarded as a sequence which has at least 70%, preferably 80 or 90% amino acid homology with the polypeptides encoded by he nucleic acid molecules according to the invention.

Nucleic acids and polypeptides which are particularly preferred are those comprising the sequences of nucleotides illustrated in Figures 1 and 2. These sequences are specific to Candida albicans with no functionally related sequences in other prokaryotic or eukaryotic organism as yet identified from the respective genomic databases.

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Nucleotide sequences according to the invention are particularly advantageous for selective therapeutic targets for treating Candida albicans associated infections. For example, an antisense nucleic acid capable of binding to the nucleic acid sequence illustrated in any of Figures 1 to 61 may be used to selectively inhibit expression of the corresponding polypeptides, leading to impaired growth of the Candida albicans with reductions of associated illnesses or diseases. The antisense nucleic acid corresponding to the sequences identified in Figures 1 and 2 may therefore be particularly useful in selective treatment of Candida albicans associated infection.

The nucleic acid molecule or the polypeptide according to the invention may be used as a medicament, or in the preparation of a medicament, for treating diseases or conditions associated with Candida albicans infection.

Advantageously, the nucleic acid molecule or the polypeptide according to the invention may be provided in a pharmaceutical composition together with a pharmaceutically acceptable carrier, diluent or excipient therefor.

Antibodies to the protein or polypeptide of the present invention may, advantageously, be prepared by techniques which are known in the art. For example, polyclonal antibodies may be prepared by inoculating a host animal, such as a mouse, with the polypeptide according to the invention or an epitope thereof and recovering immune serum. Monoclonal antibodies may be prepared according to known techniques such as described by Kohler R. and Milstein C., Nature (1975) 256, 495-497.

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Antibodies according to the invention may also be used in a method of detecting for the presence of a polypeptide according to the invention, which method comprises reacting the antibody with a sample and identifying any protein bound to said antibody. A kit may also be provided for performing said method which comprises an antibody according to the invention and means for reacting the antibody with said sample.

Proteins which interact with the polypeptide of the invention may be identified by investigating protein-protein interactions using the two-hybrid vector system first proposed by Chien et al (1991).

This technique is based on functional reconstitution in vivo of a transcription factor which activates a reporter gene. More particularly the technique comprises providing an appropriate host cell with a DNA construct comprising a reporter gene under the control of a promoter regulated by a transcription factor having a DNA binding domain and an activating domain, expressing in the host cell a first hybrid DNA sequence encoding a first fusion of a fragment or all of a nucleic acid sequence according to the invention and either said DNA binding domain or said activating domain of the transcription factor, expressing in the host at least one second hybrid DNA sequence, such as

a library or the like, encoding putative binding proteins to be investigated together with the DNA binding or activating domain of the transcription factor which is not incorporated in the first fusion; detecting any binding of the proteins to be investigated with a protein according to the invention by detecting for the presence of any reporter gene product in the host cell; optionally isolating second hybrid DNA sequences encoding the binding protein.

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An example of such a technique utilises the GAL4 protein in yeast. GAL4 is a transcriptional activator of galactose metabolism in yeast and has a separate domain for binding to activators upstream of the galactose metabolising genes as well as a protein binding domain. Nucleotide vectors may be constructed, one of which comprises the nucleotide residues encoding the DNA binding domain of GAL4. These binding domain residues may be fused to a known protein encoding sequence, such as for example the nucleic acids according to the invention. The other vector comprises the residues encoding the protein binding domain of GAL4. These residues are fused to residues encoding a test protein. Any interaction between polypeptides encoded by the nucleic acid according to the invention and the protein to be tested leads to transcriptional activation of a reporter molecule in a GAL-4 transcription deficient yeast cell into which the vectors have been transformed. Preferably, a reporter molecule such as B-galactosidase is activated upon restoration of transcription of the yeast galactose metabolism genes.

Further provided by the present invention is one or more Candida albicans cells comprising an induced mutation in the DNA sequence encoding the polypeptide according to the invention.

A further aspect of the invention provides a method of identifying compounds which selectively inhibit expression of polypeptides expressed from the nucleotides sequences illustrated in any of Figures 1 to 61 and which are critical for growth and survival of Candida albicans, which method comprises (a) contacting a compound to be tested with one or more Candida albicans cells having a mutation in a nucleic acid molecule according to the invention which mutation results in overexpression or underexpression of said polypeptides in addition to one or more wild type Candida cells, (b) monitoring the growth and/or activity of said mutated cell compared to said wild type wherein differential growth or activity of said one or more mutated Candida cells provides an indication of selective action of said compound on said polypeptide or another polypeptide in the same or a parallel pathway.

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Compounds identifiable or identified using the method according to the invention, may advantageously be used as a medicament, or in the preparation of a medicament to treat diseases or conditions associated with Candida albicans infection. These compounds may also advantageously be included in a pharmaceutical composition together with a pharmaceutically acceptable carrier, diluent or excipient therefor.

A further aspect of the invention provides a method of identifying DNA sequences from a cell or organism which DNA encodes polypeptides which are critical for growth or survival, which method comprises (a) preparing a cDNA or genomic library from said cell or organism in a suitable expression vector which vector is such that it can either integrate into the genome in said cell or that it permits transcription of antisense RNA from the nucleotide

sequences in said cDNA or genomic library, (b) selecting transformants exhibiting impaired growth and determining the nucleotide sequence of the cDNA or genomic sequence from the library included in the vector from said transformant. Preferably, the cell or organism may be any yeast or filamentous fungi, such as, for example, Saccharomyces cervisiae, Saccharomyces pombe or Candida albicans.

A further aspect of the invention provides a pharmaceutical composition comprising a compound according to the invention together with a pharmaceutically acceptable carrier, diluent or excipient therefor.

The present invention may be more clearly understood with reference to the accompanying example, which is purely exemplary, with reference to the accompanying drawings, wherein

Figures 1 & 2:

are nucleotide sequences of previously unknown function isolated from Candida albicans and which sequences are not present in the public domain.

25 Figures 3 to 35:

are nucleotide sequences of previously unknown function isolated from Candida albicans and which sequences are partially or fully present in the public domain.

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Figures 36 to 40: are nucleotide sequences isolated from Candida albicans and which have an identified function based on sequence homology with proteins

from other organisms and which sequences are not present in the public domain.

5 F	Figures 41 to 61:	are nucleotide sequences having an identified function based on
		sequence homology comparisons from other organisms and which
		sequences are fully or partially
10		present in the public domain.

Figure 62: is a diagrammatic representation of plasmid pGAL1PNiST-1.

15 Figure 63: is a nucleotide sequence of plasmid pGAL1PNiST-1 of Figure 62.

Figure 64: is a diagrammatic representation of plasmid pGAL1PSiST-1.

Figure 65: is a nucleotide sequence of plasmid pGAL1PSiST-1 of Figure 64.

Figures 66 to 106: are amino acid sequences of the appropriately corresponding DNA sequences illustrated in Figures 1 to 61.

30 Example 1

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Identification of novel drug targets in C.

albicans by anti-sense and disruptive integration

The principle of the approach is based on the fact that when a particular C. albicans mRNA is inhibited by producing the complementary anti-sense

RNA, the corresponding protein will decrease. If this protein is critical for growth or survival, the cell producing the anti-sense RNA will grow more slowly or will die.

Since anti-sense inhibition occurs at mRNA level, the gene copy number is irrelevant, thus allowing applications of the strategy even in diploid organisms.

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Anti-sense RNA is endogenously produced from an integrative or episomal plasmid with an inducible promoter; induction of the promoter leads to the production of a RNA encoded by the insert of the plasmid. This insert will differ from one plasmid to another in the library. The inserts will be derived from genomic DNA fragments or from cDNA to cover-to the extent possible- the entire genome.

The vector is a proprietary vector allowing integration by homologous recombination at either the homologous insert or promoter sequence in the Candida genome. After introducing plasmids from cDNA or genomic libraries into C. albicans, transformants are screened for impaired growth after promoter (& thus anti-sense) induction in the presence of lithium acetate. Lithium acetate prolongs the G1 phase and thus allows anti-sense to act during a prolonged period of time during the cell cycle. Transformants which show impaired growth in both induced and non-induced media, thus showing a growth defect due to integrative disruption, are selected as well.

Transformants showing impaired growth are supposed to contain plasmids which produce anti-sense RNA to mRNAs critical for growth or survival. Growth is monitored by measuring growth-curves over a period of time in a device (Bioscreen Analyzer, Labsystems) which allows simultaneous measurement of growth-curves

of 200 transformants.

Subsequently plasmids can be recovered from the transformants and the sequence of their inserts determined, thus revealing which mRNA they inhibit. In order to be able to recover the genomic or cDNA insert which has integrated into the Candida genome, genomic DNA is isolated, cut with an enzyme which cuts only once into the library vector (and estimated approx. every 4096 bp in the genome) and religated. PCR with primers flanking the insert will yield (partial) genomic or cDNA inserts as PCR fragments which can directly be sequenced. This PCR analysis (on ligation reaction) will also show us how many integrations occurred. Alternatively the ligation reaction is transformed to E. coli and PCR analysis is performed on colonies or on plasmid DNA derived thereof.

This method is employed for a genome wide search for novel *C*. albicans genes which are important for growth or survival.

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Materials & Methods

Construction of pGal1PNiST-1

The backbone of the pGAL1PNiST-1 vector (integrative anti-sense SfiI-NotI vector) is pGEM11Zf(+) (Promega Inc.). First, the CaMAL2 EcoRI/SalI promoter fragment from pDBV50 (D.H. Brown et al.) was ligated into EcoRI/SalI-opened pGEM11Zf(+) resulting in the intermediate construct pGEMMAL2P-1. Into the latter (MscI/CIP) the CaURA3 selection marker was cloned as a Eco47III/XmnI fragment derived from pRM2. The resulting pGEMMAL2P-2 vector was NotI/HindIII opened in order to accept the NotI-stuffer-SfiI cassette from pPCK1NiSCYCT-1 (EagI/HindIII fragment): pMAL2PNiST-1. Finally, the plasmid pGAL1PNiST-1 was constructed by exchanging the

Sall/Ecll36II MAL2 promoter in pMAL2PNiST-1 by the Xhol/Smal GAL1 promoter fragment derived from pRM2GAL1P.

Construction of pGal1PSiST-1

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The vector pGAL1PSiST-1 was created for cloning the small genomic DNA fragments (flanked by SfiI sites) behind the GAL1 promoter. The only difference with pGAL1PNiST-1 is that the hIFNß (stuffer fragment) insert fragment in pGAL1PSiST-1 is flanked by two SfiI sites in stead of a SfiI and a NotI site as in pGAL1PNiST-1. To construct pGAL1PSiST-1 the EcoRI-HindIII fragment, containing hIFN\$ flanked by a SfiI and a NotI site, of pMAL2pHiET-3 (unpublished) was exchanged by the EcoRI-HindIII fragment, containing hIFNB flanked by two SfiI sites, from YCp50S-S (an E. coli / S. cerevisiae shuttle vector derived from the plasmid YCp50, which is deposited in the ATCC collection (number 37419; Thrash et al., 1985); an EcoRI-HindIII fragment, containing the gene hIFNB, which is flanked by two SfiI sites, was inserted in YCp50, creating YCp50S-S), resulting into plasmid pMAL2PSiST-1. The mal2 promoter from pMAL2PSiST-1 (by a Nael-FspI digest) was further replaced by the gall promoter from pGAL1PNiST-1 (via a XhoI-SalI digest), creating the vector pGAL1PSiST-1.

Candida albicans genomic library

* Preparation of the genomic DNA fragments

A Candida albicans genomic DNA library with small DNA fragments (400 to 1,000 bp) was prepared. Genomic DNA of Candida albicans B2630 was isolated following a modified protocol of Blin and Stafford (1976). The quality of the isolated genomic DNA was checked by gel electrophoresis. Undigested DNA was located on the gel

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above the marker band of 26,282 bp. A little smear, caused by fragmentation of the DNA, was present. To obtain enrichment for genomic DNA fragments of the desired size, the genomic DNA was partially digested. Several restriction enzymes (AluI, HaeIII and RsaI; all creating blunt ends) were tried out. The appropriate digest conditions have been determined by titration of the enzyme. Enrichment of small DNA fragments was obtained with 70 units of AluI on 10 μg of genomic DNA for 20 min. T4 DNA polymerase (Boehringer) and dNTPs (Boehringer) were added to polish the DNA ends. After extraction with phenolchloroform the digest was size-fractionated on an agarose gel. The genomic DNA fragments with a length of 500 to 1,250 bp were eluted from the gel by centrifugal filtration (Zhu et al., 1985). SfiI adaptors (5' GTTGGCCTTTT) or (5' AGGCCAAC) were attached to the DNA ends (blunt) to facilitate cloning of the fragments into the vector. Therefore, a 8-mer and 11-mer oligonucleotide (comprising the SfiI site) were kinated and annealed. After ligation of these adaptors to the DNA fragments a second sizefractionation was performed on an agarose gel. DNA fragments of 400 to 1150 bp were eluted from the gel by centrifugal filtration.

* Preparation of the pGAL1PSiST-1 vector fragment
The small genomic DNA fragments were cloned after
the GAL1 promoter in the vector pGAL1PSIST-1. Qiagenpurified pGAL1PSiST-1 plasmid DNA was digested with
SfiI and the largest vector fragment eluted from the
gel by centrifugal filtration (Zhu et al., 1985).
Ligation with a control DNA fragment, flanked by SfiI
sites, was performed as a control. The ligation mix
was electroporated to MC1061 E. coli cells. Plasmid
DNA of 24 clones was analyzed. In all cases the

control fragment was inserted in the pGAL1PSiST-1 vector fragment.

* Upscaling

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All genomic DNA fragments (450 ng) were ligated into the pGAL1PSiST-1 vector (20 ng). After electroporation at 2500V, $40\mu\mathrm{F}$ circa 400,000 clones were obtained. These clones were pooled into three groups and stored as glycerol slants. Also Qiagen-purified DNA was prepared from these clones. A clone analysis showed an average insert length of 600 bp and a percentage of 91 for clones with an insert. The size of the library corresponds to 5 times the diploid genome. The genomic DNA inserts are sense or antisense orientated in the vector.

Candida albicans cDNA library

Total RNA was extracted from Candida albicans B2630 grown on respectively minimal (SD) and rich (YPD) medium as described by Chirgwin et al in Sambrook et al. mRNA was prepared from total RNA using the Invitrogen Fast Track procedure.

First strand cDNA is synthesised with the Superscript Reverse Transcriptase (BRL) and with an oligo dT-NotI Primer adapter. After second strand synthesis, cDNA is polished with Klenow enzyme and purified over a Sephacryl S-400 spun column. Phosphorylated SfiI adapters are then ligated to the cDNA, followed by digestion with the NotI restriction enzyme. The SfiI/NotI cDNA is then purified and sized on a Biogel column A150M.

First fraction contains approximately 38,720 clones by transformation, the second fraction only 1540 clones. Clone analysis:

35 Fr. I: 22/24 inserts, $16 \ge 1000$ bp, $4 \ge 2000$ bp,

average size: 1500 bp.

Fr. II: 9/12 inserts, 3 ≥ 1000 bp, average size: 960
bp cDNA was ligated in a NotI/SfiI opened pGAL1PNiST-1
vector (anti-sense)

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Candida transformation

The host strain used for transformation is a C. albicans ura3 mutant, CAI-4, which contains a deletion in orotidine-5'-phosphate decarboxylase and was obtained from William Fonzi, Georgetown University (Fonzi and Irwin). CAI-4 was transformed with the above described cDNA library or genomic library using the Pichia spheroplast module (Invitrogen). Resulting transformants were plated on minimal medium supplemented with glucose (SD, 0.67% or 1.34% Yeast Nitrogen base w/o amino acids + 2% glucose) plates and incubated for 2-3 days at 30°C.

Screening for mutants

Starter cultures were set up by inoculating each colony in 1 ml SD medium and incubating overnight at 30°C and 300 rpm. Cell densities were determined using a Coulter counter (Coulter 21; Coulter electronics limited). 250.000 cells/ml were inoculated in 1 ml SD medium and cultures were incubated for 24 hours at 30°C and 300 rpm. Cultures were washed in minimal medium without glucose (S) and the pellet resuspended in 650 μ l S medium. 8 μ l of this culture is used for inoculating 400 μ l cultures in a Honeywell-100 plate (Bioscreen analyzer; Labsystems). Each transformant was grown during three days in S medium containing LiAc; pH 6.0, with 2% glucose/2% maltose or 2% galactose/2% maltose respectively while shaking every 3 minutes for 20 seconds. Optical densities were measured every hour during three consecutive days and

growth curves were generated (Bioscreen analyzer; Labsystems).

Growth curves of transformants grown in respectively anti-sense non-inducing (glucose/maltose) and inducing (galactose/maltose) medium are compared and those transformants showing impaired growth upon anti-sense induction are selected for further analysis. Transformants showing impaired growth by virtue of integration into a critical gene are also selected.

Isolation of genomic or cDNA inserts

Putatively interesting transformants are grown in 1.5 ml SD overnight and genomic DNA is isolated using the Nucleon MI Yeast kit (Clontech). Concentration of genomic DNA is estimated by analyzing a sample on an agarose gel.

20 ng of genomic DNA is digested for three hours with an enzyme that cuts uniquely in the library vector (SacI for the genomic library; PstI for the cDNA library) and treated with RNAse. Samples are phenol/chloroform extracted and precipitated using NaOAc/ethanol.

The resulting pellet is resuspended in 500 μ l ligation mixture (1 x ligation buffer and 4 units of T4 DNA ligase; both from Boehringer) and incubated overnight at 16°C.

After denaturation (20 min 65°C), purification (phenol/chloroform extraction) and precipitation (NaOAc/ethanol) the pellet is resuspended in 10 μ l MilliQ (Millipore) water.

PCR analysis

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Inverse PCR is performed on 1 μ l of the precipitated ligation reaction using library vector

specific primers (oligo23 5' TGC-AGC-TCG-ACC-TCG-ACT-G 3' and oligo25 5' GCG-TGA-ATG-TAA-GCG-TGA-C 3' for the genomic library; 3pGALNistPCR primer :5'TGAGCAGCTCGCCGTCGCGC 3' and 5pGALNistPCR primer: 5'GAGTTATACCCTGCAGCTCGAC 3' for the cDNA library; both from Eurogentec) for 30 cycles each consisting of (a) 1 min at 95 °C, (b) 1 min at 57 °C, and (c) 3 min at 72 °C. In the reaction mixture 2.5 units of Taq polymerase (Boehringer) with TaqStart antibody (Clontech) (1:1) were used, and the final concentrations were 0.2 μ M of each primer, 3 mM MgCl2 (Perkin Elmer Cetus) and 200 μ M dNTPs (Perkin Elmer Cetus). PCR was performed in a Robocycler (Stratagene).

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Sequence determination

Resulting PCR products were purified using PCR purification kit (Qiagen) and were quantified by comparison of band intensity on EtBr stained agarose gel with the intensity of DNA marker bands. The amount of PCR product (expressed in ng) used in the sequencing reaction is calculated as the length of the PCR product in basepairs divided by 10. Sequencing reactions were performed using the ABI Prism BigDye Terminator Cycle Sequencing Ready Reaction Kit according to the instructions of the manufacturer (PE Applied Biosystems, Foster City, CA) except for the following modifications.

The total reaction volume was reduced to 15 μ l. Reaction volume of individual reagents were changed accordingly. 6.0 μ l Terminator Ready Reaction Mix was replaced by a mixture of 3.0 μ l Terminator Ready Reaction Mix + 3.0 μ l Half Term (GENPAK Limited, Brighton, UK). After cycle sequencing, reaction mixtures were purified over Sephadex G50 columns

prepared on Multiscreen HV opaque microtiter plates (Millipore, Molsheim, Fr) and were dried in a speedVac. Reaction products were resuspended in 3 μ l loading buffer. Following denaturation for 2 min at 95°C, 1 μ l of sample was applied on a 5% Long Ranger Gel (36 cm well-to-read) prepared from Singel Packs according to the supplier's instructions (FMC BioProducts, Rockland, ME). Samples were run for 7 hours 2X run on a ABI 377XL DNA sequencer. Data collection version 2.0 and Sequence analysis version 3.0 (for basecalling) software packages are from PE Applied Biosystems. Resulting sequence text files were copied onto a server for further analysis.

15 Sequence analysis

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Nucleotide sequences were imported in the VectorNTI software package (InforMax Inc, North Bethesda, MD, USA), and the vector and insert regions of the sequences were identified. Sequence similarity 20 searches against public and commercial sequence databases were performed with the BLAST software package (Altschul et al., 1990) version 1.4. Both the original nucleotide sequence and the six-frame conceptual translations of the insert region were used 25 as query sequences. The used public databases were the EMBL nucleotide sequence database (Stoesser et al., 1998), the SWISS-PROT protein sequence database and its supplement TrEMBL (Bairoch and Apweiler, 1998), and the ALCES Candida albicans sequence database (Stanford University, University of Minnesota). The commercial sequence databases used were the LifeSeq® human and PathoSeq™ microbial genomic databases (Incyte Pharmaceuticals Inc., Palo Alto, CA, USA), and the GENESEQ patent sequence database (Derwent, London, UK). Three major results were obtained on the basis of

the sequence similarity searches: function, novelty, and specificity. A putative function was deduced on the basis of the similarity with sequences with a known function, the novelty was based on the absence or presence of the sequences in public databases, and the specificity was based on the similarity with vertebrate homologues.

Methods

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Blastx of the nucleic acid sequences against the appropriate protein databases: Swiss-Prot for clones of which the complete sequence is present in the public domain, and paorfp (PathoSeq^{IM}) for clones of which the complete sequences is not present in the public domain.

The protein to which the translated nucleic acid sequence corresponds to is used as a starting point. The differences between this protein and our translated nucleic acid sequences are marked with a double line and annotated above the protein sequence. The following symbols are used:

a one-letter amino acid code or the ambiguity code X is used if our translated nucleic acid sequence has another amino acid on a certain position,

the stop codon sign *is used if our translated nucleic acid sequence has a stop codon on a certain position,

The letters fs (frame shift) are used if a frame shift occurs in our translated nucleic acid sequence, and another reading frame is used,

the words ambiguity or ambiguities are used if a part of our translated nucleic acid sequence is present in the proteins, but not visible in the alignments of the blast results,

The phrase missing sequence is used if the

translated nucleic acid sequence does not comprise that part of the protein.

Blastx: compares the six-frame conceptual translation products of a nucleotide query sequence (both strands) against a protein sequence database.

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Screening for compounds modulating expression of polypeptides critical for growth and survival of C. albicans

The method proposed is based on observations (Sandbaken et al., 1990; Hinnebusch and Liebman 1991; Ribogene PCT WO 95/11969, 1995) suggesting that underexpression or overexpression of any component of a process (e.g. translation) could lead to altered sensitivity to an inhibitor of a relevant step in that process. Such an inhibitor should be more potent against a cell limited by a deficiency in the macromolecule catalyzing that step and/or less potent against a cell containing an excess of that macromolecule, as compared to the wild type (WT) cell.

Mutant yeast strains, for example, have shown that some steps of translation are sensitive to the stoichiometry of macromolecules involved. (Sandbaken et al.). Such strains are more sensitive to compounds which specifically perturb translation (by acting on a component that participates in translation) but are equally sensitive to compounds with other mechanisms of action.

This method thus not only provides a means to identify whether a test compound perturbs a certain process but also an indication of the site at which it exerts its effect. The component which is present in altered form or amount in a cell whose growth is affected by a test compound is potentially the site of action of the test compound.

The assay to be set up involves measurement of growth of an isogenic strain which has been modified only in a certain specific allele, relative to a wild type (WT) C. albicans strain, in the presence of R-compounds. Strains can be ones in which the expression of a specific essential protein is impaired upon induction of anti-sense or strains which carry disruptions in an essential gene. An in silico approach to finding novel essential genes in C. albicans will be performed. A number of essential genes identified in this way will be disrupted (in one allele) and the resulting strains can be used for comparative growth screening.

Assay for High Throughput screening for drugs $35~\mu l$ minimal medium (S medium + 2% galactose + 2% maltose) is transferred in a transparent flat-bottomed 96 well plate using an automated pipetting system (Multidrop, Labsystems). A 96-channel pipettor (Hydra, Robbins Scientific) transfers 2.5 μl of R-compound at 10^{-3} M in DMSO from a stock plate into the assay plate.

The selected C. albicans strains (mutant and parent (CAI-4) strain) are stored as glycerol stocks (15%) at -70°C. The strains are streaked out on selective plates (SD medium) and incubated for two days at 30°C. For the parent strain, CAI-4, the medium is always supplemented with 20 µg/ml uridine. A single colony is scooped up and resuspended in 1 ml minimal medium (S medium + 2% galactose + 2% maltose). Cells are incubated at 30°C for 8 hours while shaking at 250 rpm. A 10 ml culture is inoculated at 250.000 cells/ml. Cultures are incubated at 30°C for 24 hours while shaking at 250 rpm. Cells are counted in Coulter counter and the final culture (S medium + 2% medium + 2%

galactose + 2% maltose) is inoculated at 20.000 to 50.000 cells/ml. Cultures are grown at 30°C while shaking at 250 rpm until a final OD of 0.24 (+/- 0.04) 6nM is reached.

200 μ l of this yeast suspension is added to all wells of MW96 plates containing R-compounds in a 450 μ l total volume. MW96 plates are incubated (static) at 30°C for 48 hours.

Optical densities are measured after 48 hours.

Test growth is expressed as a percentage of positive control growth for both mutant (x) and wild type (y) strains. The ratio (x/y) of these derived variables is calculated.

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References

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Claims

1. A nucleic acid molecule encoding a polypeptide which is critical for survival and growth of the yeast Candida albicans and which nucleic acid molecule comprises any of the sequences of nucleotides illustrated in Figure 1, 2, 4 to 7, 9 to 11, 13, 15 to 20, 22 to 26, 28 to 32, 34 to 43, 45a and b, 47 to 49, 51, 52, 53 to 57, 59 and 60.

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- 2. A nucleic acid molecule encoding a polypeptide which is critical for survival and growth of the yeast Candida albicans and which nucleic acid molecule comprises any of the sequences of nucleotides illustrated in Figure 1, 2, 36, 37a, 38, 39 and 40 and fragments or derivatives of said nucleic acid molecules
- 3. A nucleic acid molecule according to claim 1 or 2 which is mRNA.
 - 4. A nucleic acid molecule according to claim 1 or 2 which is DNA.
- 25 5. A nucleic acid molecule according to claim 4 which is cDNA.
- 6. A nucleic acid molecule capable of hybridising to the molecules according to any of claims 1 to 5 or the sequences illustrated in any of Figures 1 to 61 under high stringency conditions.
 - 7. A polypeptide encoded by the nucleic acid molecule according to any of claims 1 to 6 or the sequence illustrated in any of Figures 1 to 61.

- 8. An expression vector comprising a nucleic acid molecule according to claim 4 or 5.
- 9. An expression vector according to claim 85 which comprises an inducible promoter.
 - 10. An expression vector according to claim 8 or9 which comprises a sequence encoding a reportermolecule.

- 11. A nucleic acid molecule according to any of claims 1 to 6 or the nucleotide sequences illustrated in Figure 1 to 61 for use as a medicament.
- 12. Use of a nucleic acid molecule according to any of claims 1 to 5 or the sequences illustrated in Figure 1 to 61 in the preparation of a medicament for treating Candida albicans associated diseases.
- 20 13. A polypeptide according to claim 7 for use as a medicament.
- 14. Use of a polypeptide according to claim 7 in the preparation of a medicament for treating Candida25 albicans associated infections.
 - 15. A pharmaceutical composition comprising a nucleic acid molecule according to any of claims 1 to 6 or a polypeptide according to claim 7 together with a pharmaceutically acceptable carrier diluent or excipient therefor.
- 16. A Candida albicans cell comprising an induced mutation in the DNA sequence encoding the polypeptide according to claim 7.

- 17. A method of identifying compounds which selectively modulate expression of polypeptides which are crucial for growth and survival of *Candida albicans*, which method comprises:
 - (a) contacting a compound to be tested with one or more Candida albicans cells having a mutation in a nucleic acid molecule according to any of claims 1 to 5 which mutation results in overexpression or underexpression of said polypeptides in addition to contacting one or more wild type Candida albicans cells with said compound,
 - (b) monitoring the growth and/or activity of said mutated cell compared to said wild type; wherein differential growth or activity of said one or more mutated Candida cells is indicative of selective action of said compound on a polypeptide or another polypeptide in the same or a parallel pathway.
- 18. A compound identifiable according to the method of claim 17.
- 25 19. A compound according to claim 18 for use as a medicament.
- 20. Use of a compound according to claim 18 in the preparation of a medicament for treating Candida albicans associated diseases.
 - 21. A pharmaceutical composition comprising a compound according to claim 18 together with a pharmaceutically acceptable carrier, diluent or excipient therefor.

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- 22. A method of identifying DNA sequences from a cell or organism which DNA encodes polypeptides which are critical for growth or survival of said cell or organism, which method comprises:
- (a) preparing a cDNA or genomic library from said cell or organism in a suitable expression vector which vector is such that it can either integrate into the genome in said cell or that it permits transcription of antisense RNA from the nucleotide sequences in said cDNA or genomic library.

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- (b) selecting transformants exhibiting impaired growth and determining the nucleotide sequence of the cDNA or genomic sequence from the library included in the vector from said transformant.
- 23. A method according to claim 22 wherein said cell or organism is a yeast or filamentous fungi.
- 24. A method according to claim 22 or 23 wherein said cell or organism is any of Saccharomyces cervisiae, Saccharomyces pombe or Candida albicans.
- 25 25. Plasmid pGAL1PSiST-1 having the sequence of nucleotides illustrated in Figure 63.
 - 26. Plasmid pGAL1PNiST-1 having the sequence of nucleotides illustrated in Figure 65.
 - 27. An antibody capable of binding to a polypeptide according to claim 7.
- 28. An oligonucleotide comprising a fragment of from 15 to 50 contiguous nucleic acid sequences of a

nucleic acid molecule according to any of claims 1 to 6.

Sequences with unknown function, C. albicans sequence NOT present in the public domain (ALCES/EMBL)

>328c2 1803bp in-house: 1123-1803 public: 1-436/468-1021 PathoSeq: 437-467/1022-1122

ATGTCTATTACAGTTACATTTCCGAAATCTCCATCTACGAAAAAACGTGCACCGGCATTTGGAATTGAGTTGGAGTTYAG

TCAMCAAGSCAGTAGCGATGGTGCTATAGAGAAAAGCGGCATTGGCAGTTCCT GTGTTTAGCGTTGACAACCAAGACTWT

GTATTKATAAGAGAYCWTGCCAAGTACTGGGGCTACCCTTCATCGTATCAATT GATTGTCAAGTTGGTCAAATGTGCTAA

CATTGAAAAGTCGCAAATCTTAAAGACCGATAAGGATTTGAATAGAGAGTTGT TTGAGTTGGATTTGAATAGAAGCAG

ATACAAAGATTGATCTTTTTATATTTCGTTACCCTTGGTCTATTCAAGAATAGA AAATAAGAAGGTTTTTTATGTTCTG

CGTGAACCAGAACAGCCAAAGGTGTCGAAAGCMCCAACACAAGAGAAACCAG CAAGTGTGGTTGCTGCAGAAGAAGATGA

CGATAATCTAGATGATGAGGAGGAGGACGAAGTGGATGAAGACATGGATGAA GATAATGATAATAGTGGGGAATTGTCTA

AAGGATACAAGCACATGCACAAGGACCATCCAAAGTATATAAATGACGATAGGGTTACTATTGGACAAGTGTTTCATCAA

TACGGACTTGACCCTTCGACACCATTAACCCATTCACTTTTCAATAGTATCAACTCAATGTCGAAGCTAAACTATTACAA

GAATTTTGGAGTTTCCGATTTCTTCCCAACAGCAAGTTATCTTATGC AGAACGAGAATTGGTGTTGAATGCCA

ACAACTACAATGATATGCACATTAACGAAAAGACAGAATCCAAGCCGAAAAA GAGTTTCCGTAAACCCATTGGAAAGTCA

AAGAAACATAACTTGCAGATTGATCCGAACTCCATAGATTTAAGCGAGTCAGT GATTCCGGGACAAGGGTTTATACCTGA

CTTTAGTATCCACCTATCTTTGCAAAGTCCCTAATTATTATGTGACATCAACCC ACCAAAGTCTCCCGCTGTCGTTCAAC

ACAAAGAATCTTAATGCAACTTCGAACTCTTCGTATTTGTTTAATGATAATGTC AAGATAAAGTCAAAAAGTATTCAGAA

GTWSGTGTTCAACAGCGATACCGATAATTACCATCACACAAAGTATTTCTACA CCAAAACCTACCGTGGTCCAGGGTCGG

GGAATTACAAGGATGGTGCATTGATGAACAAAATCAACAAGATACATCTTTCC AGTAATAAAAAGCCGCGCCACAAGAGA

AAGGTGTCGAACAATAACAGGTACAACAAGAGTTTAAAGGGGTTAGTCCACG AAAAGTTTGACAAGAACTTTGTTGAGTA

CTTGCTTTCTGAGCAACGCAAGTATACCGAGGACTATTCCAATCTTGAAATTTT ACACAATAGCTTACAGTTTAATGTTC

TTTTGAATACGTATCGTGGTGTTGCCCAAGAGACATGGAATAACTACTACAAG
TTTAAATTGATTGATTTCGAACAATTG

AAGGCTTTGCAAATGGAGGCAAATGAGCTTGAGGAGAAAATTGGATGCTG CTAGACACCAACAGTGGGCGGAAGAAGA

GAAGCTTTNCCAAGAAAGATTGCGTTTAGTATTTGAAGATGAACGGACGAGTT TGAGCAATTGCAAAGCGAGTTTGGTCA

GAGAAAGAAGATTTGGAAGAGAAATTGCGTCGCCGTCAGCTANANGCATCTT TGANTGATAGTTTTGAACTTGATAGCG AAAATGACNATGAATCTTGACTTGNCCAAANTNAACAAGACTT F19 (6.4)

>214c cpL1 290bp inhouse:1-290

Fig 2

Sequences with unknown function, C. albicans sequence present in the public domain (ALCES)

>113g2 638bp in-house: 1-638

CTTATTCGGTTCTAGTGTCTCAATTGGTTATCCATTAACATCTATTCCCAACTCC ATCATTATTGGCAATAAATAAATGG GTGTTATATCTATTGGTAATAACTAAACTGGTGTCAATTCAATTCCAATATGGT CATGACAATTGAAAGTGTTACTGTTC TGGTTTACATATTCTACAGGTTACAACTATTGATTGGTTAGAAGTTTGGTTTCA ACATCACCTGTTGCTAAGAATAAATG TATCTATTATCTACAACCACCAAGTG ATAAATGCTGAACCGTAGTCACCAACTGTTATGCTGGTTGTATCTATTGACTAA AACTACCCTAGGGATAAATGCTGAAC CGTGGTTACCAACTGTTATGCTGGTTGTATCTATTAACTGCAACCACCAAATGA TAAATGCTGAACCATAATTACCAACT GTTACATTGCTGGTACTACATTAAGAATAAATGCTGCATCTACAAGTACCACCT **GTTGTGTTAATAAATGCTGCACCTGC** TAGTACAACTGTTGCTGGTCATGATAGTTACTACACACTTACACACCAGACAGTG **GCAAACAAGGTTATGTAGAAACCA**

>117c_af 623bp in-house: 1-623

AACTGTCCTGTGAAGACGAACATCACCACAATCATGGTCATAACCAAAAT CACAATCATGTTGCTCCTATTCCTACA ACAGCTGGACAATCATTAAATAATAAAATTGATACATCTAAAGTGACAGCTCT CAACATGGCCAACTCTGCTGACGATCT AGCAAAAGTTTTCAAAGATTCGACTAAAAAATATCAAATCAAACCAATTATCA AATCAGACAGTGATGAACAAATGATTA TCAACATTCCATTTCTTAATGGTAGTGTCAAATTGTATTCGATAATTCTACGTAC CAATGGGGATTTGTATTGTCCCAAA ACAATAAAATTATTCAAAAATGACACATCAATTGATTTTGATAATGTGGATTCG **AAGAAACCAATACAGGTGTTAACTCA** TCCTCAAGTTGGTGTTGCTAATAATGATAGCGATGATCTTCCAGAGTTTTTGGA ATCAAATAACGATGACGATTTTGTCG AACATTATGTGTCTCGACATAAATTCACTGGGGTAAATCAATTGACAATATTTA TTGAAGATATTTATGATGAANGAGAA GAAGAGTGTCATTTACATTCAATTGAATTGAGAAGGGGAATTCACTGAATTAA ACAAAGACCC F19 5

F19 6

- 1 QQSYVPQSQP NYSQQTCDRG MFSGGGGGHG HYQQQQGYNA YGPPPPQGGY
- 51 YOOOPGGGG YYOGGOCOOP MYYOOOPRSG GNDSCINGCI AALCYCCTID
- 101 MLF

>17g1 731bp in-house:1-604 public:605-731

GCTGTAGTTTTGCTTCCAAAAGTTTGATCTCGTCATCAACATCATTAACTTCATC TAATAAGGTGAATAATTTGGMTCCM KGYTCCACGTGSYYGYATCACTTTTWATAGATTTCACTTYGGACAAWACTTTA TTTCTYYGYYGATCCCATTTCYKGAMA GATCCGTGTAATGTTTCKGCGYMAGACATGTCTTTATTATATMGTTCATTTAAA GAATAGTGACTCTCTGACAACTGATC AAAGGTCKGTARAATCCTACTTCGTAATTGATATATGATTATTACCACTCTG TAGAAACTTGCCAKATTTGACTGAAT CTTCGTATAACTCTKGTGTGAGCRAKTTTACTCTGTTAGATAAATACTCGATTG GTGAKKGTGAAKTGTTGTCMTTTGAC TGGTACRKGGCTGCRGGRARAKKGATRGATTTKATCATCMAACTGTCCATGGT ATTRKRTAACAGTKCACTTYCTTTGAT AGAATCAATTAAAGTTGTGGTAGTCACTAGATTGGGTTTATGATTGTCGAGTAA GTGTGATAGTTGCTCATCACTTATAT GCTTATCCAGAAATTTATTGTACAGCACCATGTGAGTCTTTTGTAGCTGGTTTA TATACTTTATTTAANATGAACTCTTC GGGATCGAGTTCATCTTCATCTTCGGAGGTGGAAGCGGGGATAGAATGTAAAC GTTTGATAGGGGTGTCTTCTTCAT **AAGGTCCCAGA**

Fig 8

>222g8 543bp in-house: 143-543 public: 1-142

CTACAAAATGAACAATAGACAAGTTCCTAGTAGAACACCACTAGACACCAACA CTAATCCATGTACTTTCTCATTGGTAG TACCACTTTCAAATATCCAATCAAACACATCAAATTTGATTGTACTGTTCTTAT TTTTGAATGAGCTCAGNTGNNTTCTT CTGCTACCATAAAAGAATTTGGGAATTTCAAATATTGTACTTTCAAAAGNNGAT AATGCACTAGTAGAAGTGCCAAATAC TGTTGTAGTAGGTTTCTTTGTCTTTGGTGTATTAACAGATATTACTTTATTGTCT TCAATATTGATTTGTTCAAAAGGAT CCTCAAAATTTGGCGTCTTCGTGCCTAATTCTGAATTGGATAATAGTATTTGAG CCGTAATCTCATCACTATCGNCTTCT TTACCATCATCTTCGTACCTATAATAGTAATAATAATNGTCAGACAATTCAGTT TCACCACTATNAATTTCAGAATCCGT TNCGTCAACAAGATTTTTTAAAATAAAATTGTCAAACATTGACGNTGCAGTAG NGGTTGGAAA $F_{19} \quad 9 \quad (c \sim 4)$

>222g9 804bp in-house: 1-575 public: 576-704 PathoSeq: 705-804

TTCCAGAGGCAACAAGCGAAGAAGCACAACGAAAGAAGGAATTTGAACAAA AGGCCGAATTTATCATTNGCATCATTAC TTGAAATGCGCCGAAGAGAAATAGAGAGGCGGAAACAGCAAAAGGAAAGGG AACAAAGACAAAAGGAGCACGAAGCAAAG AGGGATATCAGGATACAACAACTTTCAGAGCAGGATTCACGGAGTAATCAAAC TAAAGAAGAAGAGGAAGTGTTCAAGAA GGCCCGGTCTACTAATTCGGGAGCAGACGAGACTGGTTTGATGTCAGATAAAG AGTTTGATGATTCTGCATATTCACCCG ATTATTTGTTTGAAGAGAATTTGTGGAATAAACCAAATCATCCAGATACAAATC ATAAAACCAAAAAATATACTGAGAAT GTGGTTGAAAATCTAGATTCTCCACCAAATGATACATCTGCGTACAATTCAAGT TTTCATGATGAAACTAATATTCAAAA TGAGATCCAAATACCAGAAAATGACGAGTATGTACCACAGATGAAAGCTACAT CCAGTGTCAATAATACCACCATCCCTG CACAAAGAAGACATGAGTCACTTTCCACTTCTGAAAACAAAAGAAGGAAATTT GAAACAGCCGACGTTGGGGTTGATGGG TTAGATTCTCCAGTGCGGGCACAACCAGAAATATCTGCAAAATCCAAGTCTCC GATAATCCCTGGATGGTATACTTTTTT ATGGNACCNAAGAGACTGGAAACTCCTGAAGGCAAATTGCTGTGCAGGGACC AATAGGTACATTATATTCCCTCANGGGG **GNCC**

Fig 12

>22g3 (5') 535bp in-house: 1-535

>22g3 (3') 426bp in-house: 1-426

>24gG 522bp in-house: 1-522

>28gK 475bp in-house: 1-475

>328c1 681bp in-house: 1-681

TGGTGATTTAGCACAAAATGAGAGATTAAGTTTAGAAAATGGNTGGACAGGNA TTGGGNCTTTGGNAACATTATGAAAAT GGGGATCGGTCCAATTGAGAAATATTGCTCCTGCTGGTGGA F19 17(con+)

>33gK part1 1171bp in-house: 1-588 public: 805-1171 PathoSeq: 589-804

TCTAAAATCTTCAGTNCCATNCAGAGTTTTAGGGAATGTTACAGACTCAACTCC TTTTGCCATGGGGACATTAGGTTCAA CATTTTATGCTGTCACTTCTGTTGKCWGATCTTTCCAAATYTAKGASKKGGYW ACATTACATTTATTGTTTGTTTCCCAA ACTCAAACTCCTTCAAGGAATWACATGTTTGGCTGCACACCATCAMTTTGTCT **ATGCATCTKATGGTGATCGTATTGGTA** TTTTTAGACGTGGTAGATTAGAGCATGAATTGGTTTGTGAAGGGAACTCTACAG TYAACCAATTATTAGTATTTGGAGAA TACCTTATTGCTACCACATTAGAAGGTGATATTTTCGTATTTAGAAAAACTGAA GGAAAGAAATTCCCMMCTGAATTATA CMCTACMATCAGAATAATTAWTYCTTTAGTTGAAGGAGAAATTGTGGGATTA ATTCMTCCACCTACGTATTTAAWWARWR TWRTTGYTSCWMCYACYSWWTCTGTGTTTGTTATAAATGTGAGAACTGGCAA ATTATTATACAAATCCCGGGAATTACAA TTCGAAGGCGAAAAGATTTCATCAATCGAAGCTGCTCCAGTTTTGGATGTAATT GCTGTTGGTACATCTAATGGAAATGT ATTTTTATTCAACATTAAAAAGRGGGAAAGTGTTGGSCCAAAAAATTATTACTT CTGGAACTGAATCTTCCTTCGAAAGT TGCCTCGATCTCYTTTAGAACAGATGGAGCACCTCATTTGGTTGCTGGTTTGAA TAACGGAGACTTATATTTCTACGATT TAGACAAGAAATCACGTGTTCATGCTTTGAGAAATGCCCATAAAGAGATTCAT **GGGGGTGTTGCAAACGCCAGATTTTTG** AATGGTCAACCAATAGTATTATCAAATGGTGGTGATAATCATTTGAAAGAATTT **GTTTTTGATCATAATTTAACCAGTTC** GAATTCATCCATTGTTCCTCCTCCAAGACATCTCAGATATAGAGGTGGGCATTC AGCACCACCAGTAGGTATAGAATTTC GTCAAGAAGATAAAACCCATTTTTTATTGAGTGCTTGTAGAGATAAAACATTTT **GGACATTCTACTTTGAGAAAAGATGC**

>33gK_part2 1001bp PathoSeq: 1-1001

TCAAGCACAGGAATTGTGTCAAAGATTGCAAAAATCTAAGGATGTATAAAG

GGATGATTTGTCCATAGTTGTTATTGACGTGACTACTCAAAAAGTCATAAGAAT ATTATATGGTCATACCAACAGAATTT

CAGGAATGGATTTCTCGCCTGATGGGAGATGGATAGTTTCAGTTGCATTGGACTCCACTTTGCGAACTTGGGACTTGCCA

ACTGGTGGTTGTATTGATGGGGTGATTTTACCAATTGTGGCAACTGCAGTTAAA
TTTTCTCCTATTGGTGATATCTTAGC

GACAACACATGTCTCTGGAAATGGTGTATCCTTATGGACTAATCGTGCCCAGTTCAAGCCTGTGTCCACCAGACACGTAG

AAGAAGATGAGTTTTCAACTATTTTATTACCAAATGCTTCTGGAGATGGCGGTTCAACAATGCTAGACGGGTTTTTGGAC

GAGGATTCTAATGAAGACGCACTATTGATGAACAGTATACATCTGCTCAAATTGATGCATCCTTGATTACTTTATC

ATCAGAGCCAAGATCAAAATTCAACACTTTATTGCATTTGGATACCATTAAACAACAAAGCAAACCGAAAGAAGCACCTA

AAAAACCAGAAAATGCACCTTTCTTTTTACAATTGACTGGA Fig 18 (cont)

>33gK_part3 414bp PathoSeq: 1-414

AAATTGCGTAAATTGGATACAAACGGTAACCACGCATTTGAAAGTGAATTCAC AAAACTATTAAGGGAAGCTGGAGAGAG

TGGACAATTTGAAAGATTTTTGACTTACTTAACTTATCTCCTGCTGTATTG GACTTGGAAATTAGATCACTTAATT

CATTTGTTCCATTGACTGAAATGACAAATTTTATTCAAGCTTTAAATGCTGGTTT GAAATCAAACGCAAATTATGAAATA

TGGGAAACTTTATATGCCATGTTTTTCAACATACATGGTGATGTTATCCATCAG
TTTGAAAAATGAAACTAGTCTTCATGA

AGCTTTGGAAGAATACAGACAGTTAAATGATGAAAAGAATAACAAAATGGATT
CTTTAGTGAAATATTGTGCTAGTATCG

FIG 18 (conf)

>35gK 1334bp in-house: 146-669 public: 1-145 PathoSeq: 670-1334

ACAACGTATAATCGACAGTTTACTATATCTGCTGACTTCAAAACCAATGCATTC
TTCAAGCGTGCTCTGTCGATTTCTAT

CATAACATCCACTTTCCGGNGTAATCGGATTACTAAAGCCACAGAATCAAGGT GAACATCAAGCTTCAACTTCTTTCTTG

GTCCACGAATAATTTTAATTTGGTTMTTSKKGSMAMKGCTTTCTACRGTAGGTT TGAATCTTTCCAACATTGTCTTTGCA

TAGAAACMGCACCAGACAAGAAACATGTCCACTCGACCATCAACYTSKGGGT AWWGACAAAGTWAATCTGTCTGGATCCT

TITCATCCAGTTTCCCTGCATKGGAWACAAGTNTGTCCCGCACAGTTAAGACT GTTTTTATTTTSKTGGTATTAGACTCA

TCAAGTTCCGAAGGAGGCATCATTTARGGGWATAGACTCCGCTGAGTTAAT ACTGGATAAATCACTTATTTCAGATTC

ACTGACTTGTWCTTCAGTGACCTTATCAAAATCCTCAATGTACTCSGARGCGTW TTCMCTCMATGTGAAGGCTTTTAAAA

GGGCAACRCTGGTTYCAAAATGCTTTCTTGCRAGTTTGTACKTGACAGAAAAA TCAAAAACYTTGAAAGATATACCTCTT

F19 19

CTAAAGTCTTTTAAATCAATTTCTTNTCCTAATTTTTCATCATATAGCTTATGAC TTGGCAAACCCTCCTTACATACCAT ATCCATTACAATGCTAGAAATGTCAATCTTCACTGACGATATAAAGGATGGAA GAACTTCAAATAATTTTATAAACTCAG GATTGGCTGGTGTATCTGCTGCAGGAGCTCCAGATTTATTGTCCATTTGCTCAC TCCATGGACATACATTATTAACGTCC ATCTTTTTCCATTCTTCAAATTTCTTCGGTGAAATAAATTCGTTGACGRWTTTTA AACAGACGTACAATGTGAAAGATAA TTTTGTTGTAACATATTGTAGGTGGCT AAAAGATTGACTTWRGTAAAATGRAACTTATTAACCCTGGGCCCTCACATTTC ACATTTTCATCTTAAACAAAGKGGTT CAAAGKGGAACTTGGTTTGGATCCYTTAWTGGAAWATTTCYCAGKRAATACTT TCAAAATCAACTCCAGGAGAGCCACAG TGATAATTGAATTGGATTTAGATAAGCGGTTAAACTTCCCAATTTCAGTTTTAC CAAACTCTGGTAAATGAAGGTTAAGT TTTGTGTCCACCACAACAAGTTTACTAAAAACAGCCTTGAGCATTTTGGAGGCA Fig 19 (wat) >36g2 (5') 520bp in-house: 1-520

>36g2 (3') 472bp in-house: 1-472

>38g1 1348bp in-house: 183-940 PathoSeq: 1-182 / 941-1348

GATTTTTTTCTGGTGCGCCGGACACGCCCTCCGGTCCGCACCGAAAACCGGGG TAATCTCCGTCGGAGATACACATCCG

CGGACACAAATCAGATGAGCTACCACCGAAAATTCCGAAATTTCAAAAACTC AAAATCCCTAAAAACAAACTATCCAGA

NATTATTGCCATGCCCTGAGGATGAGTTTAGTTTTTAATTTTTGAAAAATGTCCAAAACTGGTTGTGCTGTATAGGANG

GGTAAGAATTTGCCATTCTGCCCCTTTGGGTGGGTCAGTCNAAAAAAGANGTA TCACTCTGGTTCNAACGGGAAACAACN

NAAAATGGGATTAAAMTWATCTCCAGAMCAAACTTAGCTTMWWACACCCAY TTTAGTTGTACTSGYGWRCCMAAMMCMAA

TTTTCCATTTTGTTTGGGGANGGGAATTTARACCAAAWTTTTTTTTTTGAAATTT CGCTMAGTGTYMAGAMCCSCAAAAG

TCACCTTTTTCGTTTTCMMCYACGGCARARGCYCACCGGTTTTKYKTGGKGS MCRGCCMAATTGAWTTTGTGGGTGSGC

ACGKGGAAAAACAGTTKGTTAGTGGACACGTTTTTGCAGTGTGAAACTGCGCTCGGAGGTACTATATGCGAAAGCAGAAA

AGACAATTGCAAGAATACAGAGAGTTCTTCTCTGGGCTANNGCAATGTGTTTA AGGCCAAGTCGACGAGTGGGGAGAGTC

TGGAAGTGATATACACATCACGACCTACTTTATACGCTACGTTCGGCATGGGC GAGCCACTGTACGGTGGCAAGCCTGAA

CAGTCCCACACCAGATATCTAACGATTCTGTGTATGGGCACTGATGGGATTTAG TGGATTACTAGCTGATAGCAAGTATT

GAAAACTAAAACCCGACTCGGGGGTATGCCTTGGCAAGTAGCCGGAGTAAAAT CTGTGACTTTGCTGAGTGTAACTCCCT

CCATGGTTGGCGATGTTCGACGTGCGCGGCAGTTCTTGTCGTATCACAGTCGCACGGACACCACACCGGGAGAATCTTAA

GAGGGCTATATGGATGTGGAACGGTTTGCTTGCTGTGGTAAAACACTGGCGGG CGAGCCGACGTTCCACGGACACAGCAA

TGTGTTTGCAACCAAATAAATAACTTGTACGGTTTGAACGTGTTTTTGGCTGCT CCTTCCAGTTCTTGGCGGGAGAAGCT

TGGGCGCGGGAAGACCACTACTACGTAGTTATCTGGTTGATCCTGCCAGTAGT
CATATGCTTGTCTCA

F19 21

>3gG 842bp in-house: 171-842 public: 1-170

ACCANTAAGGGANGAGGTGGACAAATTGAACCAGAATTGGGAGCCGCCAAATTGGACGGAAACCCCGGGGAGGCGGCCAA

TTTGGGCGAGAACCCACGGATGGACAAGCGGGGCAAGTGCCAAGGAAGTTGCGTGCCAGTGAAGTGGAAATGTTGGAGAGC

TGGCAATGAACTGCCCGGCAAGTGATACCAGGAGATAGGTGTGTATAGATTAT
NATGGAACGCCNATTTTTGCAGTATCA

CGCGTAATAAGGACAGCAGTTGGACATCGGTACATGAGAGAGCAATGTNAGTC TTGATANTAATGAGCCGTGTTGAAGTA

GTATTTTAATCNAATTTTACTCCCAAAAGGACAATGGAGATCTGGAGATAACN CCACACTAATCGGTTCTAGACATAGAC F19 22

>480c 731bp in-house: 1-731

TTTGAANTCTTCNCCNCCTGNNNCTNCTCAAAGCGCTCCGCNTTGTGNCTNAAN **GGGCTGGCTCAACGTACTATCAAGNG** TAACATAAAACTANNACTTGGGNCATTCACGATAAAGAAACGGNGCCTGGAAT TCCAGCAATTGNCGCTGGACTTGAGTC GNAATCTAGGNATTGNTAGNCGAGATTTTNCATATGAANNACCTGGGAAACGG ATCAACTGGCTTAACANGACCAGTATT GATGNGGAGAAAAGTGGGACTTGCAGANTTTCTCAATANCCTCATTCAAGA CTCAACACTTCAGAATGAACGAGAAGT GTTGTCGNTTTTGCAATTGCCGNCTAATTTTAGATTCACCAAGGATATGTTACA GAATAATCGAGCNNACTTGGATTNTG NGCAAAATAACTGGTACGATGTATATCGTAAGTTGAAACTGGATATACTCAAC GAATNGTCTAGCAGCATTAGTGNACAG ATACATATTCGTGATCGCATTAGNCGGGTCTACCAACCACGGATTCTCGACTAG GCAGGNCTATTGGTACAGATAAAGAA GANGCCTAAAGAAGAAGCAGNTGGGTTCCCAATTCTTTNAGAGNNTAGAAAAT TTGGTAGTACAGGAAGTTTCCCGATCA AAGAGGGTGTTGGACCCAACAGNTTAANNTTTCGGCNGNGNCNTTNTCATTNA ACAATAAGNACTTCTTNAACACCAAAT Fig 23 CCNNGTTTTTA

>55g3 1063bp in-house: 533-1014 public: 88-532 PathoSeq: 1-87/1015-1063

TTCNCCCCCATTCCAAGATTCCCCTTGTAAGTAAATTGGTTGGAGAACCNCGTT
GGTTTAATTCCCCCGNNCGGGAAANN
ATTNGTNGTAAGACAATTCTTTCAACAATTTATGATGTTGCTGCATTTTCGTAA
ATCTGCTAGTAAGAGCACCCTAATCT
TCGAAATCAATAGAAAACTCAATCTTTTCGATTTTCTTGTGGATCACTTTACTC
ATAGGCTCGCTATAAACAAACAAGCT
CACTATGATAAACATATAAGGTAAAGTATTGACTTGATGTAGATGTAATCACCA
CAGAAGGCATATCAATACCAAGTCGAA
GAGGAGCCTGTTTCGAATTATCATCATTGTAATTCGCTTGCTCTTCTTGATCCA
GATCTTGTGCATTTGGAGAAAGCTTG
CTTGACAAAATTTCACTTTCATAACACATTGTCATAACAGAAAGCTTTTCAATC
AATAAGTTGTTGGCTCCAGCCCATTT

ACCATTTTGGGTTATTTCTGTTCCTAGCCATGGTGGCCAATTAGATTTAGCTCC **ATATGGATTACTAATACTTAACATAT** CTGGACACCCTACAATATCCTCTTTGTTTAATACAAAAACATTGGCATCTTTTA **GTAAAATACCATACCTAGTTTCTAAC** AAGATTTGGTTTGCATTATTCCTGGAATCCCTAATGGACAAAATTTTACCTTTA ATAGATGGTGTAGAGATGAGCACNAC AGAATCAGGATAATCCTCNCTTTGTTATTGAATTTGAGGTGGGANCACNTCNAT CAAATAATCCTCAACCACTTTTTCGT CNGGTCTGGTCTCNCNTAATATTGTGTTGAAATTGTCCAGTTTCCGTTGTGAAG TGANATCNTTGCCGGTAGANGTCTGT GATTTAACCCCCTCCCGTGTGNAACGACCGAATANTTCTGACGTCCANNAAAA ANCGTCCCTATCGTTGTTTGCNTTTAC NCCATCCNCCNTTTTTCCAGTGTTTTCCATGGACTTGTGTGNCGAGTTATTTCG AAGCTGTGATTTCAATTTCACAAAA Fig 24 (con+) TGTATGTATTTCAATGTCAAATT

>58gA 724bp in-house: 281-582 public: 209-280/583-724 PathoSeq: 1-208

GTGGTGTTTTGGAAAGTAAGGTGTGATTTGCTTAACAAAGGAAAAGAACGAGA CGAGAACCAATTCTCAATATATAGGTC TTTCCAGGTAGGAAAACGACCAACTGTGGAAGAATGGCACTACCATTGGTTAC CAATACAACTACTGGTTGTCGTTTCAT CTGATAACATACACAACTACCATGTAGAAGTCATCAATTGTAAACTCAGCTAT ACTTTATCAATATCAGCGAAATTTTAT TTAGTTTTCGTTNGAAAATACAGTGAAAAATAAAAATCTACCCGTCNNCNTGA ATTGTNTNCCTCTGCANCNACAAATNG TNNTTATATTGTGATTCATTTCNNAGGCTTGATTNCCANCTATTTNTAAACACT ACCTTTCATTTNCTACNTTCNGGGAA AATAACNCTTGTTGCTGTTGAAAGACCAATNCCNTTGTGAGTACAGAGGAATA CTNCCANTATNCGGCTTATANTTANCT AAAATTACAATACATAACAGGGAACCAGACNTGTTTNCGTCNTTGATAATGAA CAGTTNTGGTNCTNNTGAAAAGTAATC CCNAATTTGAATGGNTCGAAAGCAACACAATAAGAGTCTTTGCTTGATATTTG CTTCTCCAGAATATAAAATAACGCGTT

F19 25

>60gK 990bp in-house: 445-752 public: 1-140/753-990 PathoSeq: 141-444

TAGATCCTGTTTTTTTTTTCTTCGCCACA

TCAC

ATTACCGATCCGTCGGATTTTAAAACCACAAAATTGCCTGCATTAGCAGAGCT
AGATATTTTCATAGGGTGCTATATATG
CAAAGATCTATTGAATGCACCCGTGAGGACACAATGTGATCACACGTACTGTT
CACAATGTATACGAGAATTTTTACTTC
GAGATAATAGATGTCCGCTTTGTAAAACAGAGGTTTTTGAAAGTGGTCTAAAA
CGTGATCCATTGTTAGAAGAGATCGTC
ATTAGTTATGCCTCCCTTAGGCCTCATTGATTACGATTATTGGAGATTGAAAAG
GTGGAATCGAAGCAAGAGGTAGATCG

Fig 26

TGAGAAATCAGCCAATGAGTCAGCGCTGAATGGTAATAGAAATGTAAACAAC GATGTTGACGAAACTGTGCGCGTTAAAG

ATCAACTGAATGCAGATAAACTAGGTGAAGAAAAAGGGCAAGCTCAACATGGGGAACAAGTNAAACGAGCAGACTACTGA

AGTTATTCTGTTGCTATCTGATGATGAAGAGAATGGTTCTGATAGCCTAGTAAA ATGTCCTATTTGTTTTGAGAGAATGG

AATTAGATGTACTACAGGGAAAGCNTATTGACGACTGTCTAAGTGGAAAGAGC ACGAAGAGGCCTACAGACATTTTA

CCAGTTNNGGCGTCNACAACTCCCACAGCAACTCCGACAACTACATTGTTGAA AGCAAACGTCTCATCTCCATCCCAAGT

AGTTGAGTGATATGAAACTACCAACAACAGGTAGTAGGAATGAAATGGAAGC CAGATACTAGCATTACTATGTGATTTAT

AATGCCAACCTTGACACCAATCATCCTGTA

Fig 26 (cont)

>61gB 602bp in-house: 1-602

ACCTACNNTCACNCNNGNCNCNGCAACACCANCNTNCCNCCNAAANAAAGTC TTCTTTGAATNAGACNTTTCATCTATTG

TNGCCGTNGTNGTTGTTTCTGAGCAATGAATTCTTGTTGGATTTGCNTGNTGTC ACGTTCATNTTCTTGATTGAATTTAA

TACCATTTTGAAAATCATTAAAATCTGCCACTNCNACAGTTACTCTTTTGT CTTANGATACAAATNNTACCCTTTT

NCACGTTTGGCGTTTGTGNCCAATGCAAACNCNGTNCCCCGGGGNGNGGGGNCCCCCCCCGCCNTTGTCCANANCCTGNT

GGNTAGTTATNTTCGCGCTTACNATCCCCCCCACCCNNGCGNGNNCCGCCCCA
CCCCCAACGTNCNCCTCCTCTCCCCNC

NNCTNGCNGCGCCTNTCNGGGCACCCCTTCNCCNCCTCNC Fig 27

>62gB 539bp in-house: 101-539 public: 1-100

ATAGAATCAGAAACAAAAGCTKTAGTTTGKGAAGAAATTGAAACAATCGGA AAACAACAATATCAAACTGMTGCCCAAT

AACACTGGTATGTACCTAGATGGATTACCAAGATCTACTACATAAAATAATAR RGGAGTTCCACTCACTCAAAGAGTTCA

AACCATGGGATAGCAGTGTTTTGTATGAGACGTTACTACGATCAGTATTAACTACTTTTGATCGAACTTTTGGGCMTAGAC

AATCCACCCAGTTWTCTWCMCCTCACCACCMACMATGATAGTTATAGGTGAA
TTTGAAAATWAAATACTATGGRAATGCA \mathcal{F}_{19} 28

TTAAGCAAGTCAATCAAYSSWCMKRGCATRKTGCAATWTGCYWGWAYCAWA GSATGYAATCSATATTACMRRCYKTKSYY GRGRYYRKKRATACGCGAWMATWTWKAATCMMMGAGTCYWATYCTGCTGK WWTCMAAGA F19 28 (cm+)

>64gB 627bp in-house: 1-627

TNCANCCTNCCATNCNCCCAGGCNNNGCCACCCCNGCCGNNCCCCCNTNTTTC CCCCCTCTTNGTNGCCCTCNNGGTG GTGTTTGTGGTGTGACNAATAAANATGGTNTATCATTAGAANAGGACATTGCN NCGGAAATGACTGTCGACAATAAAGAA GCAAATATATACAATGGATTATGAANGTGCTAGGATGGATTTGAAAGTTTATC TGGGTTTATTCCAATGTAAAAATTATT TGTAATTGATATGGCTAATTATTTTGCTCNATATNTATCACAAAAAAAATGATTA **AGTTCGAAATGAAATTGGCNTCCATA** TATAAAATTTCTGACAGGAAGAGAAAATTCANGACNTGTTGCCCNAAAAAAA **AACTTTACCCCNCNTCNANTCNTGTNN** GACTTAACCCCCAAAAANAANANNGCTGGCGGCGGNAAAAAAATAGGAGGGG GCCGGNNGTTTTTTAAAATTTNANNCTT GAATATGAACCCAANNTTTGNNTTCNTTTTTNCCACNCCCCCTTCAAATTTNAT TCCATGTTCCCAAGANNAGGGNGGNG GGGGNGGTTCCNNCTTTTAAACCNCCCCCCGGGTGGNGGGGNCCGTNTTNT TTCCGGNGGGGCNT F19 29

>65g 441bp in-house: 1-441

TINCTTATGTAGATGTTGTTCATGAATTTGTATGAACGGACTATGGCTAGGATT
TGGCCAATCTCGGTATTACTACNTTT
TCAAGTTCAAAGATTGGGAAACTCGTGTATTTTCGTACTGTCTACATTTTCTTA
AATTTGATAAACGCATAGTAAGTCTT
TGCTTGATATACTATGAGATGATTAGAATTAAAAAAGTAGACGACTAGTTTCACT
AGATTTATTGAAGTGTCAAAATATAT
TCAGATTGGTTGCAACTGATGGTCTCGAAAATGCNACAGGATTTTTTTCCCCCA
TITTTTGCCAATTTTTGTCCNATAGA
GTAGAAAGTACCNGTATNCNAATTGTCCCAAAAAAGCGATTATAATCCGTACCA
ATATTTCCAATTTTCNTTTAAACCCTG
TTCNCCTCNGTGTTGGTTTGTTTTGAAAACNTAACCANGGTG

FIG 30

>8c_cp 890bp in-house: 287-890 public: 1-124/154-286 PathoSeq: 125-153

ATGCAATTCTCATCCGGTGTCGTCTTATCCGCTGTTGCTGGGTCCGCTTTGGCTG
CTTACTCCAACTCCACTGTTACTGG
CATTCAAACCACTGTGTCACCATCACTTCATGTGAAGAAAACAAATGTCACGG
AAACTGGAAGGTTACCACTGGTGTTAC
CACCGTCACTGAAGTTGACACCACCTACTGCCCATTGTCAACCAC
TGAAGCTCCAGCTCCATCTACTGCTA
CTGATGTTTCTACCACCGTTGTCACCATCACCTCATGTGAAGAAGACAAATGTC
ATGAAACCGCTGTCACCACCGGTGTC

F1931

ACCACTGTCACTGAAGGTACTACCATCTACACTACCTACTGCCCATTGCCATCT ACTGAAGCTCCAGGTCCAGCTCCATC TACTGCTGAAGAATCTAAACCAGCTGAATCTTCCCCAGTTCCAACCACCGCTGC TGAATCTTCCCCAGCTAAAACTACTG CTGCTGAATCTTCCCCAGCTCAAGAAACCACTCCAAAGACCGTTGCTGCAAT CTTCTTCAGCTGAAACTACTGCTCCA GCTGTCTCTACCGCTGAAGCCGGTGCTGCTAACGCTGTCCCAGTTGCTGCT GGTTTGTTGGCTTTGGCTGCTTTGTT CCCTTTCCCTTTCTTCATTCTTCAAA AAAGGGTTATTTACTATTAATTGATAAATTTATGGTTTCATGTTAATTTACCCTT TTCTTTATAAACATTGGTATTATTA TTATCATCATTAGNTTTATTTATATTTTCGTGAGTTTTTCGGNTTTAATTAATTTT TTTGGATACATATTAAAAATTTAT Fig 31 (cont) TTGGTACTAG

>80g3 669bp in-house: 1-652 PathoSeq: 653-669

TTGCCAAAATTTTATAAAAAATTGTCAAATTGAAAAGAAGTATTTCCCAAAAT **AAATTGTTTTTTCATCACAACCGGTTC** ATATCGCCATAGNCCATTTTTAATCTTAAGGTTGATACCAGTTAATTGTTGATTT CTCTGTTATAGCCCCTGTCTAAATC TGTCTATTTCTGGTATCGAATCAAAATGTCGCTCATAATGTGCATGTCGCAAAG ATGTCGTAAAGTTTTGATTTCATACT CATCTTAAATTTTTTTTTAGTGATTGGCATTTTGTTCTTTCACATAGTTTTTATTTC TAGTTATCAACCTATCAAATACAC CTCCACAACAATGCATCCAAATAATAAAAATTCATTTAAATCAAAAAAGAAAT TTATAGATCGTCGAGAAGCCAAGTCTC AAACTATTAGAAAAAGAAGGGTTACAA GGAATAAATTTTGAAGAACGTGCAGCCAT TGTGAAACAACGTAAAGAGGAAAAACGTAAATTCAAACTAGCAAGTGTACAA GCAAAATTGGAAAAGATTGAATCTAATT CGAAAGAAGAGCTTTAAACCGTGACCAC

>21g2 667bp in-house: 1-667

TTAGAATGAGTAAATTGAATGGAAA ATCACTGCAACAACAACAACCACTGGTGGATACGAAAATTTAGTGTACAA ATTTCTGCCAAAAAAATACAATAAAAA CCGCTTATAGTCTTCTACTGACATAACAACACAAGTCAATAAATCAACAACTC **ATAAACAATGTAGACTTAATACTATCG** CTTAATTATTTAAACTATAATAAATACCCTATAGTATTATGCCTTTGTCAATGTG TGTAGAATTTGGTTATTACATATCC ATGTGTNATATATGTTGATCAAAAAAACGCGATCTTCTCTTTGGTGTAGTGT **GTTACNCAAAAAATTCACTANTCTAG** GTCNCTGANAATCACTTGAAAAATCAAAAATTTGTTGAAATTTGAATTTCCTCMA YTTTGAAATTTTGTTTGAAATTTTTTT TTTGCTTTACAAAAAGACTCCATTTTGTTTTCCATTTCACAACCAATTACTTAAT TCCTCTTTTCATAATTAATAACTA TCATTACTTACAACTACAAACAACTACGATCATTTCCTAAGAAAAAGCAACGA GGGCGAATTGAGACATTAATCCCCTTT F19 34 ATTTTATCATCATGCCTTATACAGAAC

>66g4 579bp in-house: 1-579

Sequences with known function, C. albicans sequence NOT present in the public domain (ALCES/EMBL)

>CFL (223c cp) 165bp in-house: 1-165

AACTATTGCCAATGGTAAATATGCCAGTGAAATCGAGAATTTTAATAAGTCGG
TCCCTCTTAAGGTCCCATTCAAATTCA
CTAATGCACAATTGGATCTTTATGCTGCTAGCACACATAACCAAGAGCCAATA
TCCTAGTAACGACGCACCATAGTAGAC
CGAAT

36

>EF4 (29g3) part1 479bp in-house: 130-479 PathoSeq: 1-129

CGCGAAGNNTCAATCATNTCAGAAGAAGAAGAAGGTACTCCGTTCTTTAC
TATTGTGGCAAGAATCCCTGTGATTGA

GGCATTTGGGTTTTCCGAGGATATTAGAAAGAAGACATCCGGGGCAGCTAGTC CTCAATTAGTTTTTGATGGGTATGATA

TGTTAGATATCGATCCATTTTGGGTTCCACATACTGAAGAAGAATTAGAAGAAT TGGGTGAATTTGCAGAAAGAGAAAAT

GTTGCTAGAAGATATATGAATAATATCAGAAGAAGAAAAGGGTTATTTGTTGA TGAGAAAGTCGTCNAAAATGCTGAAAA

GCNAAGAACTTTGAAAAGAGATTAGATTATCCNGTTNAACAGGCCATATGTGT GAAATTGTTTCCNAAAAGACAGATACN

ANGTGGNCCNTATTTGTTTAATATTCCACNACCAGTTAATGTTTTGATATNGAT GTTTTATATAGTCCAATGTTGAGAC F_{19} 37 (c)

>EF4 (29g3) part2 1706bp

AAGTCATGCGATTGCAACAAGGATCACAAGAACCAGAAGTTCACGAACATTTG
ATTAATTTGATTGATTCACCTGGGCAT

ATTGACTTTTCGTCTGAAGTGAGTACTTCTTCGAGATTATGTGATGGTGCAGTTGTTTTTGGTCGATGTCGTCGAAGGTGT

CTGCTCACAAACAGTCAACGTTCTACGCCAATGTTGGATTGATAAGTTGAAGC CATTACTAGTTATTAACAAAATTGATA

GGTTAATCACAGAATGGAAATTGTCTCCCTTGGAGGCATACCAACACATTTCC AGAATTATAGAACAAGTAAACTCTGTG

ATTGGGTCATTTTTTGCTGGTGATAGACTAGAAGATGACTTGAATTGGCGTGAGGCTGGTTCTGTCGGGGAGTTTATCGA

GAAGAGTGATGAAGACTTGTATTTCACACCTGAAAAGAATAATGTAATATTTG CCTCGGCAATAGATGGATGGGCATTTT

CAGTCAATACATTTGCCAAAATATACCTGAAAAAATTAGGGTTCTCTCAACAA GCATTGTCAAAAACTCTCTGGGGAGAC

TTTTACTTGGATATGAAAAATAAAAAATCATCCCTGGTAAAAAATTGAAAAA
TAATAGTAACAGTTTGAAGCCATTATT

TGTTTCGTTGATTTTGGACCAGGTTTGGGCTGTTTATGAAAACTGTGTTATTGA AAGAAATCAAGACAAGTTGGAAAAAA

TCATTGAGAAATTAGGGGCCAAAATCACCCCTCGTGATTTGCGATCCAAAGAT TACAAGAACTTGCTAAACTTGATTATG F_{ig} 37 (5)

TCTCAGTGGATTCCTTTGAGTCATGCCATATTGGGGTCAGTGATTGAATACTTG CCAAGCCCCATTGTTGCTCAGCGTGA

AAGAATAGACAARAWWTWRRKGRRMCSMYYTATARWRYWKTGKWTTCAR AAMTGSATAWWTCCAAACTAGTCGAMMCTT

CATTTGKMAARRMKWTGCASRMAYSYSMKRGTWMACMCCCRKAAMCCMAT WGGCMATWKYAKNTGTMYCAAAATTGTTTG

TCAATCCCCAATGAAGACTTACCCAAAGCTAGTAATGCCCGCTACTGGAGGA TTGACGCCGATGAAATCCAAGAACGA

GGAAGAATTGCTCGAGAATTAGCCAAAAAGGCATCTGAAGCAGCTGCTTTGGC ACAAAGAAGGTTCCCAAAAATGAAGAT

GAGTTTGCCATTAAACCCARGAAAGATCCATTTGAATGGGAATTTGAGGAGGA CGATTTTGAGAATGAGGAAGATGAGAG

CGATGCAAACGCAGTTGAAGAATCAACTGAAACCATAGTGGGTTTCACTCGTA TTTATTCTGGATCGTTATCTAGAGGCC

AAAAGCTCACGGTAATTGGACCCAAATACGACCCTTCATTACNTAGAGACCAT CAAACCAACTTTGAACAAATAACCAGT

GAAGTGGAAATTAAAGATTTGTTTTATATCATGGGAGGAGAATTAGTGAGAAT GGAAAATTTCCGTGCGGGTAATATTGT

TGGGGTTGTTGGATTGGATAACGCCGTGTTTAAGAATGCCACAATTTGCTCACCGTAACGTGAAGATAAACCATACATTA

ATTTAGTTTCAACATCAACCTTGATCCACAATAAACCAATTATGAAAATAGCA
GTTGAACCAACAAACCCAATAAAACTA
GCAAAATTGGAACGAGGATTAGATTT

Fg 3765) (cont)

>NDI (17c_cp) 807bp in-house: 1-614 PathoSeq: 615-807

AACCTATTCCATAATGTTTACTAGATCATTGATTAAAGGTGGTGGCAGACTTGC TACTACCAGATCATTGGTCAACAACT

CTACTAGTTTGGTTTTAAAAAATCAATTTAAGAAATATTCAACATCAACTCCTC
CTAAGGTTGCCAAATCAAAATCTTCG

ACAATTGGTAAAATATTCAGATACACTTTTTACACTGCTGTGATATCGGTTATT GGTTCTGCCGGTTTGATCGGTTACAA

TTATTTTGGGTTCTGGTTGGGGTGCTATTTCATTATTGAAAAACTTGGATACCA CCTTGTATAATGTTGNTATTGTCTCC

CCAAGAAACTATTTCCTTTTCACCCCATTGTTACCATCTGTTCCTACCGGTACTG TTGAATTGAGATCTATTATTGAACC

TGTCAGATCAGCAGAAGATGCCCTGGCAAGTTATTTACCTTGAAGCAGA AGCTACAAATATNAACCCCTAAAACTA

ATGAGTTGACACTTAACAAAGTACTACTGTCCGTTCTGGTCATTCTGGTAAAAA
TACTTCCTCTTCTAAATCAACTGTTG

CCGAATACACTGGGGTTGAAGAAATCACTACCACCTTGAATTATGACTATTTA GTTGTTGGTGTTGGTGCTCAAACAATN

>RPL27 (357cL) 560bp in-house: 1-560

AAAAATGGCTAAGTTCATCAAATCTGGTAAAGTTGCTATTGTTGTAAGAGGTCGTTACGCTGGTAAAAAAGTAGTCATTG

TGAAACCACATGATGAAGGTACCAAATCTCACCCATTCCCACATGCCATTGTC
GCTGGTATTGAAAGAGCTCCATTGAAG

GTTACCAAGAAGATGGATGCTAAAAAAGTTACCAAAAGAACTAAAGTCAAGCCATTTGTTAAATTAGTAAACTACAACCA

TTTAATGCCAACTAGATACTCATTGGATGTTGAATCATTCAAATCTGCTGTCAC
TTCTGAAGCTTTAGAAGAACCATCTC

AAAGAGAAGAAGCTAAAAAAAGTTGTCAAGAAGGCTTTTGAAGAAAAACATC AAGCTGGTAAGAACAAATGGTTCTTCCA

AAAATTACACTTTTAAGAAAGGAACCACCTTTATTTGAATGTTTGTAATATAGG TTGAATCAGAGAGACAAAGTAGAAGA

AAATACAAAAAGAGAGTATATCTGTATAGTATAATTTAATGGGGGTCTAATT TACTTACCACTTTATTCGTGCATTATT F_{19} 39

>SADH (110c_af) 650bp in-house: 1-650

AACCTTTTGAAACGATTAANTNCAATCAAACAATCTTATTCAAAAGTACTCGC AATACGTACAATGTCAATTCCATCTAC

TCAGTACGGATTTTTTTATAATAAAGCTAGTGGTCTTAATTTGAAAAAAGACTT GCCGGTTAACAAGCCAGGTGCTGGTC

AATTGCTTTTAAAGGTTGATGCAGTTGGCCTTTGTCATTCAGATTTACATGTTCT CTATGAAGGTTTGGATTGTGGTGAT

AATTATGTGATGGGCCACGAAATTGCTGGGACTGANGCTGAACTANGGTGAAGAGGTGAGTGAGTTTGCAGATGGAGATC

GTGTCGCTTGNGTCNGNCCCCANTGGATGTGGNCTTTGCAAACACTGTCTTACT GGTAACGATAAATGTNTGNACCAANT

CGTTTATTGGATTTGNTTTCGGATTGGNTTACAATGGNANGNTNCGANCCATTT TTGGTAGNNANGAGANCNANAACTTG

GTAAAGATCCNTNTAATGTACNTNCCGAGNAAGCTGCNCTTTTNNNGNNTCCN TATTGANTCNTACCAANGNTTTTAAGG

NATGNAGNAGTGTGNCANCNTGGGAATAATACACNTTCTTNCCTGATGGN

NTTNTGCNGTTACNCNNTTCAAGNNNN F19 40

Sequences with known function, partial C. albicans sequence present in the public domain (ALCES)

>ABP1 (409c10) 1435bp in-house: 842-1435 public: 1-382/779-841 PathoSeq: 383-778

ATGGAAAAATTGACATGAATACGTATTCAAACAATATCCAACAAGCATACGA TAAAGTTGTTAGAGGAGAACCAAATGC

AACATTCGTCGTTTATTCTGTTGACAAAGACGCCACTATGGACGTCACTGAAAC AGGGGACGGATCATTATACGATTTTG

TTGAACATTITACTGATGGACAAGTTCAATTTGGTTTACCCAGGGTTACTGTTC
CAGGATCTGACGTCTCCAAGAACATC

TTGTTAGGATGGTGTCCTGACAGTGCTCCACCAAAATTGAGATTGTCATATGAC AATAATTCTGCTGATGTGTCAAGAAT

ACTGAGCGGATACCATGTGCAAATTACTGCAATGGATCAAAATGATTTATACG TGAATCACTTCTTGAATAGAGNTGGTG

CTGCTGCTGGTGCAAGATATTCCACTCAAAACTTCCGGACTCAAAAAACCATCCCCCTGCTGCACCTAAACCTACTTCAAAA

CCTGTTGTTGCTAAATCTAGTTCTGCTTCAAAACCTTCATTTGTACCCAAATCTA CTGGGAAGCCTGTTGCTCCAGCTAA

GCCAAAACCAAAGAACATCACCAAGGATGCTGGTTGGGGTGATGCTGAAGAC GTTGAGGAAAGAGACTTTGACAAGAAAC

CTTTGGATAACGTTCCATCGGCATATAAACCAACAAAGGTTAACATTGACGAA TTGAGAAAACAAAAATCAGATACAACT

AAGGATGAAAGCCTATGATCACGACTCAAGTCGTGATGGAAGATTGACTTCTT
TACCAAAACCAACGATTGGACATTCTG

TNGCCGATAAATATAAAGCTAGTGCATCTGGGAATGGTGCTGCTCCTGCGTTTGGTGCTAAACCAGCATTTGGTCACAAT

CAGTTGATTCAAGAAAGGATAAATTGGTAGGTGGTTTGTCGAGAGATTTTGGT GCTGAAAATGGAAAAACTCCGGCACAA

GCCAGAGGAACATCATGCTGCCGACTTGGCCAAAAAATTTGAAGAAAAGGCA AATATTGCTGGCGATACTCCTTCCTTGC

GAGGAAGAACAAGCTTCAGCACCATCTTTGCTACTAGAAACTTACCACCACCGTNACAAAGACAACCTGAGCCCGACCAG

AACCAGANGAAGAGGAGGAAGAAGAAGAGGAGGCTCCTGNTTCAAGCTT ACCAGCAAGAAATCTCCCCCCAC F_{ig} 41 (Lont)

>ADE12 (226c_af2) 993bp in-house: 1-646 public: 647-677 PathoSeq: 678-993

NATAAGGAATGANCCAAAGNAGTGTANNAGNTAATAAGNTNAGANANTTCCA AAAGAANNAAAANAACCTAACNACANAN

NNNTATTANATCCAGTAGAAANACAANATTAAGNNACCANTATCTTNNAAAG
NTTNACGANACNTNGTTCNAATTGTTCA

TNTTGGGANGNACCCTCAACAACTNTANTNGGGTAAACTTTAACNACACCATA GACATTTNTNATGGTTNTTGAGGAATA

CCCAACCCAGTCANAAANACCAACAATACCAGTTGATAATAAGTGANGTATGA TAAGTACAGANATCAATATCCAACATT

AAACGCATTAGCACCTTCANCCAAGATTTTTTTATTGGCAGCNATAGCTTCGTGCATNAAGTTGACGGAGTCTGACGACG

AATGGTCTCAAGGTTTCACGGNATTTTTCAAATCTTGCCAATTCTTCCTTAGGA TCATATTCAAATTCACCGTATCTTTT

TTGTCTACTCTCGACTAATCTCAAATATCTAGTTTTGAATTCTTCCCAAGCTTCT GGATCAGGGTTGACTAAATGGTGGA

CTCTGATACCTGATCTACTTGCCTTGGTTGAGTAAGTTGGACCAATACCTTTAC
CGGTAGTACCTATTGATTTCTTATTG

GNTGTTAATTCAGCATCTTTCAATTTATCAGCACGTTGATGGAAGTCAAAGACC AAATGAGCTCTAGATGAAACAAACAA TCTATCACGACAATCTAACCCTTTTGCTTCCAAGTTTTCCAATTCAGCAAAGAA **GGAAGGAACGTGGATAACAACACCAG** ATCCAACTAAGTTTTGACATTTAGGATTGACCAAACCAGAAGGTAACATGTGG AAGTCATACTTGACTTTACCAACAACA ATCGTGTGGCCAGCATTGTTACCACCTTGACATCTGGCACAAACATCGATATCA TCACATAATAAATCGACTAATTTACC Fig 42 (con+) TTTACCTTCATCCCCCCATTGAGATCCTAATAC

>CDC48 2448bp in-house: 95-220/285-1340 public: 1-94/221-284/1341-1373/1783-2273

PathoSeq: 1374-1782/2274-2448

CGTTTCCGAWAGCTGCACCTGAGTTGGCACCTGCTGCAACCATTATCAGTG

GCACCAGCATTTTCATTGAATCTAAAG

CTAGAAAATTGACCTCTTGAGGCTTGCAATTGTTGAGCGTAAGACTCATAACG

ACGTAATTCAGCGTCTGAAACAGATCT

TTTTGCGGTCTTCATAGCCTCTTCAAAGTGAGCTCTGGTAATGTAAGGCACAGG

GTCTTCTTCTTCAACTTCATCTACCT

TCATATCAACATCTTCAGTTATCACCTGTTCCTTTTGCTTCTTTAATCTTGTTAAT

CTTTACTTGGGCTTCAATAGAGTC

TTTAATAGCAAATTTAGCAGATCTTTGAACAATATAAGACAAATCTGCACCCG

AGAAACCGTGAGTGATCTTGGCAATIT

CGTTCAAGTCCAAACCAGGTTCTAATGGAGTGTTTCTCAATTGAGCTTGTAAAA

TAGACAATCTAGCTGGCTCATCTGGC

AATGGGACATAAATTAATTGATCCAATCTACCTGGTCTCAATAATGCAGGATC

AATTTGATCTGGTCTGTTAGTGGCACC

AATGACAAACACATTCTTCTTAGCATTCATACCGTCCATTTCAGTCAACAATTG

ATTGACCACTCTGTCGGAGGCACCAC

CAGCATCACCGTGAGAACCACCTCTAGCTTTGGCAATGGAGTCCAATTCATCC

AAAAACACCACAGTAGGAGCAGCAGCT

CTGGCCTTGTCAAATATATCACGGATATTAGACTCAGATTCACCATACCACATA CTCAACAATTCTGGACCTTTGACAGA

AATGAAATTAGCAGAAACTTCAGTAGCAACAGCCTTGGCCAAAAGTGTCTTAC

CAGTACCTGGTGGACCAAAGAACAAAA

CACCTITTGTTGGTGCCAATCCGAATTTTTGGTATTGATCTGGATGTAAAACAG

GATACTCCACGGTTTCTTTTAATTCA

TTCTTAATGTTGTCCAAACCACCAATATCATCCCAAGTGACATTAACATTTTCA

ACAACAGTTTCACGCAAGGCAGATGG

GTTGGAGTTTCCGAGAGCAAATCTGAAGTTGTCTTGAGTGACACCCAAAGAGT

TCAACACTTCAGTATCAATGGTTTCTT

CTTCCAAGTCGATAAGATCCATCTTTTCACGGATTTGTTGCATAGCAGCTTCTG

AACATAATGAAGCAATATCAGCACCA

ACGAAACCATGTGTTTCAGAAGCGATGGCTTCCAAGTCAACATCATCAGCCAA

TTTCATATTCTTTGTGTGGATTCTCAA

AATCTCTAAACGTCCTTCAGCATCCGGAACACCAATGTCAACTTCTCTGTCGAA

TCTTCCATCAATAGAATTTTGAAATC

TTCTCAAAGCAGGGTCCTGTTAGTAGCAGCCAATTAACAACCTACATTAGATCT

GGCCCTTCATACCATCCATAAGGGGT

TAACAATGGAGAAACAACTCTTCTTTCTACTTCACCCATTAGTTTTGTCTCTCTT TGGGGCAATAGAGTCAATCTCATCA ATGAAAATAATGGAAGGAGAATTCTTTTCAGCCTCTTCAAAAGCTTTTCTTAAA TTGGATTCAGACTCACCAGCCATTTT AGACATAATTTCTGGACCATTTATTAAGAAAAAGAAGGCACCTGTTTCATTGG CCACTGCTCTTGCCATAATGGTTTTAC CGGTACCAGGTGGACCATACATCAAAATACCCTTTGGTGGCTTAATACCAATC GATTTGAATAATTGTGGATGTCTTAAA GGCAATTCAACCAATTCTCTAATTTGGGCCATTGTTTCTTACACCTCCCAATAT CGTCGTAACCCACTTCATTCAAGCTA TTTTCTTCATCTTCACGATTAATAGGTTCTCCTTCACAATGAATAATGGTATCTT GAGCAACAATTGCAATTTCTTCAGG GTCAACTTCAACAACTTTGAATTCTACTTGTCTCATACCACCCCTCACAGTGAA TAAATCACCTTTTCTCACTGGTCTAT AGGCTTCAACAAAATATGGCTTCAAGTAAAGGTCGAATAAGGAACCATTAATA CCTTCAACAGTATCAGCAATTGGCAAT

ACGAACACATCTGTTAACTCTAGCAACGCCATCAGGCATATCATCATCAGCTA
AAACGATCAACACTGTGTCCTTTCTCT

TCTTACCTTTCACCAAGACTGTATCACCACGGAATAATTGTAACAATTCCATTG
TGTTTGACGACATGGTTATGACAGAA

ATCTACAGCAGAAGCACCAGAAGCATCAAAATGTTGTTTTTTATCTTC Fig

Fig 43 (con+)

>CIT (99g3) part1 1435bp in-house: 803-1435 public: 1-333 PathoSeq: 334-802

TCCAAAACTTATTGCTTAGCTATACGGTGTAATGGACCAGCTTAACCATTCAAA CCAGCAGCTAATGACAAGAATGGAGA AGATAAAGCGGTACCAACCAAGTGGGTGGTGTGCAGAGACGTTACCACCTT CGTGGTCAGAGTGGATGGTAAGGTACA ATCTCATTAATTCAACANATTCCTTGGTGTCACCAAAACCTAACANACTGGCCA AGTTAGCACCGTAATCCAATTTGGAG TCAATGGCAGCTGGCAATTTACCATCGTGGAAAACGTTTCTGTAAATCTTAGCA GCAATGGTTGGCAATTTAGCTAACAA ATCGATGGAATCTTCGTAAGTGTATTTCCAGTATTCGGATTTGTTGGCACCTTT AGCATAAGCTTGGGCAAATTGGGATT CAGATTCCAAAGCAGTAACGGCAATGGAGAATTGAGCCATTGGGTGCAAGTGA GATGGAGATCTGTCGATCAATTCTTCA ACGTGCTTTGGTAATGCTGATCTAGCAGCAAATTCTTCGGATAAAGCCTTAGTT TGGGCGTCAGTTGGAACTTCACCAGT CAACAACAACCAGAAAAGAGCTTCTGGTAATGGTTCTTCACCACCTGGTGCTTT TGGCAATTCTTTTTGAATGTCTGGGA TGGTTCTTCCTGAAACGGATACCTTCAATTGGGTCCAAAACAGAACCTTCCC AAACTAAACCTTTGATACCTCTCATA CCACCGTAAGCTTGTTCTAATAAAACTTCACCCAATGACAGTTTTACCGTGTTC

Fig 44(9)

TTTTTGAATTGTTTAACTTCTTCAG

CTTTGGCTGGCAAGATTTCTTCCAATCTTTGTTTTAAGGTCTAAGAAAGCAAGT TAGTAAATTATCTTTGTATATACGTT

GCTGTTTTTGAAAGTGCTCTTGGCTACGTTGGTTGAACGTTGAATTGATCTGAA TGCAGACATTGTTTGATAAATATACT

TATATACAAAAAATCCAGCANGAAATCAGTTGAAATTTATATTCCAAATTTTT
GGTTTTAATGGCTCTTGAAGTTGTGG

TGAACAATTTTTTTTTTTTTTTCCTCATGGATTTACTTTAGTTTTGGGTC TGTCGNGCTGCCGTACAACTTTCC

TGNGAAAATTGATTTTTTTTTTCTTCTGGNGAGGATTTTTTTGGCGTTCTTTGTTAA ACTTCTTATTTGTCCCACTANATGG

>CIT (99g3) part2 327bp public: 1-327

CAAAGAGAATTTGCTCTTAAACATATGCCAGACTACGAATTGTTCAAATTGGTT
TCAAACATTTACGAAGTCGCTCCAGG
TGTTTTGACCAAACACGGTAAGACCAAGAACCCATGGCCAAATGTGGACTCCC
ACTCTGGTGTCTTGTTACAATACTACG
GTTTGACTGAACAATCTTTCTACACTGTCTTGTTCGGTGTTTCCAGAGCCTTTGG
TGTCTTGCCACAATTGATCTTGGAC
CGTGGTATCGGTATGCCAATTGAAAGACCAAAATCTTTCTCCACTGAAAAATA
CATTGAATTGGTCAAAAACATCAACAA
AGCTTAA

>HOL1 (409c5) part1 695bp in-house: 98-695 public: 1-97



>HOL1 (409c5) part2 762bp PathoSeq: 1-762

GATCAGAATAATGAGGACTTTATACCTGGAACACTCAATATCTATTCCTTGGAA GTTGACTCTGAAGATGAAAACGTGAG TCATTACGATGCTTCCAGTCGACCAAAAGTGAAAACAAAAGGCAATATAATCC TCTTCCCACAACCATCGAATTCATGCA ATGATCCATTAAATTGGAGTAAATGGAGAAAGCTAAGTAACTTTTTTATTGTCA TTTTTATTACTGCTTTTACAGCAGCT ACTTCAAATGACGCTGGATCAATTCAAGATTCACTTAATGAAAAATATGGAAT TAGTTACGACGCAATGAATACAGGGGC TCGTTATATGGTCGAAAAATAACAT AAGCACTTCCGACTCAATTTGGTCG CAATTGTTTGTTGGTATTAGTGAGAGTTGTGCTGAAGCTCAAGTACAATTAAGT TTATCAGAACTTTATTTTGCCCATAA **GGACCTTTAATTGCAGCCTTTATTG** TTCAAAACATTGGTTTTAGATGGGTTGGTTGGATTGCAGCAATTATTAGTGGTG CATTATTGTTCGTAATTGTTTTTGT TTAGATGAAACCTATTTTGATCGAGCAAAGTTTACCAAGCCA Fig 455

>ESP1 1458bp in-house: 889-1458 PathoSeq: 1-888

CGATTTTCAATTACAAGATATTTTGCATCATGTTGAAAGCAAATGGTTTGGTGG **GTTTATTTCAGGTATTTTCACTAATG** ACAATGACGTTGAAAATGAATCCAAGAACGTGTTTCATAAATTCAAACAAGAT TTAATGAAAATTTTGAAAGATTGTTTA ACCGTAAGTGACGATAAATCGAATATAGAGAGGTTTCTTCAGTTTAATGAATTT ATTTATTACTGCTTTTACTCAATGGA GGAATATAATTATGAATTGGTTGATGATTTGATAAAATTTATAACTATAAATAT GAATTCTCATGGCAGAATAGTTAATT TTGGCACTAATGTTAAAATTAATANATTACACGAATTAATTAAGAATTTGATTG ATAAAGTTAATAAAAACAAACAAAGA AACAGCAACATTCCCAACATATTGGTT TTGATACCTAATGCCAACTGNTCCAATTTCCCAATGGGAAATCGATGGAANTTC NTTCGTAAGTAAAATCCAATTTCAAG ACAAGAACAAGTTAATGTTTGTTGATA AATCTAATTTGTATTTGATTAATCCCAGTGGTGATTTAATTCGATCAGAAA ATCGATTCAAAAAACTATTTGAATCA Fig 46

AATCATTTATGGAGAGGGGAAATTGGAAAATTATCAAGTAATGAACATGAAGA TTATCAAGATTCAATATTATGTGAAAT

CTTGAAAAGTCATTTATTTGTTTATATTGGTCATGGTGGTTGTGATCAATATATT ANAGTATCAAAATTATTTAAAAAAT

TACCTCCTAGTTTATTGTTAGGTTGTTCATCAGTTAAATTAGATAATTGTAATTA TAACTATAATTCCAGTATGTTACAA

CCACTGGGTAATATTTATAATTGGTTGAACTGTAAATCGTCAATGATACTCGGG AATCTATGGGATGTTACTGATAAAGA

CATTGATATTTTTACACTTTCATTACTACAAAAATGGGGGTTAATAGATGATTA TAATGGCAGTGGCCATGATTATGGTA

TGAAGAATTGGATTTGACTAATTGTGTTGTTCAAAGTCGAAGTAAATGTACTT TGAAATACTTGAATGGATCAGCACCT

GTGGTTTATGGTCTACCAATGTATTTÁAAATAGACATTCTGTTTGCATATAAGT TTATATATTTTAATAATAAGAAAAAG

AGCATAATTTGGATCTTGATTTTGTATTGTTTGGTTTGTTATGAACAAATTTTGC ACCCAATCACTATCGAACTTTCTTT

TTTAAACAGAGAACATTTAATCAACATTTATGTTACATTTAAGCGTTTAAATAC ATATTTGTGTTAGATAGTTATATAAT F_{19} 46 (ont)

>FAL1 (190g3) 1439bp in-house: 1-770 public: 861-1299 PathoSeq: 771-860/1300-1439

ATAATGGATGATTTTGATAGAGATTTAGATAATGAGTTGGAATTTAGTCATAAA
TCAACGAAAGGAATAAAGGTTCATCG

CACTTTTGAAAGTATGAATTTGAAACCTGATCTTTTGAAAGGAATATATGCCTA TGGATTTGAAGCACCATCTGCTATTC

AATCTAGGGCTATTATGCAGATCATCAGTGGTAGAGACACAATAGCACAGGCACAATCTGGAACTGGTAAAACTGCTACT

TTTTCTATTGGTATGCTTGAGGTTATAGATACTAAATCAAAAGAGTGTCAAGCA CTTATCTTGTCTCCTACTAGAGAGTT

GGCAATTCAAATACAAAATGTGGTCATGCATTTAGGAGATTATATGAACATTC ACACCCATGCCTGTATTGGTGGGAAAA

AAAAGAAGAAATCTACAAACTAGAAATATCAAGGTTCTTATTTTAGATGAAGC TGATGAACTTTTTACAAAAGGGTTTAA

AGAACAGATCTACGAAATCTACAAACATTTACCACCTTCGGTTCAAGTAGTAG TTGTTAGTGCCACATTGCCACGTGAAG

TATTGGAGATGACAAGTAGTTTACCACTGATCCAGTGAAAATCTTGGTGAAGA GGGATGAGATTTCGCTTCTGGGAATCA

CACAATATTATGTTCAATGTGAACGTGAAGATTGGAAGTTTGATACACTATGTG
ATTTGTATGACAACCTTACAATAACT

CAAGCAGTGATATTTTGTAATACCAAATTGAAGGTGAATTGGCTTGCTGATCAA
ATGAAAAAGCAAAACTTTACTGTTGT
GGCAATGCATGGTGATATGAAACAAGATGAACGAGATTCAATTATGAACGATT
TTAGAAGGGGGAATTCAAGAGTATTAA
TATCTACAGATGTTTGGGCAAGAGGTATTGATGTCCAACAAGTCTCGTTGGTAA
TAAATTATGATTTGCCCACCGATAAG
GAAAACTATATTCATAGAATTGGACGATCAGGTAGATTTGGTAGAAAGGGAAC
AGCTATAAACTTGATAACTAAAGATGA
TGTGGTCACTTTAAAAGAATTGGAGAAATATTATTCAACGANAATTAAGGAAA
TGCCAATGAATATTAATGATATAATG

>FBP1 (40c_af) 638bp

ACATCATCACGTTGACCCGTTTTATT TTACAAGAACAGCAAACTGTTGCTCCCACCGCCACCGGTGAGTTGTCGTTGTTG TTGAATGCGCTTCAATTTGCATTCAA GTTTATTGCCCACAATATCAGAAGAGCTGAGTTGGTCAACCTTATTGGTGTTTC TGGCTCTGCCAACTCTACCGGTGATG TTCAGAAGAAATTGGATGTGATTGGTGATGAGATCTTTATCAATGCCATGAGAT CTTCCAACAACGTCAAGGTTTTGGTT TCTGAAGAGCAAGAACCTTATTGTGTTCCCAGGTGGTGGCACATATGCTGTT **TGTACTGATCCAATTGATGGGTCGTC** CAATATCGATGCTGGTGTTCTGTTGGTACGATTTTTGGTGTGTACAAGTTGCA AGAGGGGTCTACTGGTGGCATCAGCG ATGTCTTGCGTCCTGGTAAGGAGATGGTCGCTGCGGGGTACACCATGTACGGT **GCATCTGCCCATTTGGCATTGACTACA GGTCACGGNGTCAATCTTTTTACTTTGGATCTCANATGGGTGAATTTATTTTGC** CNATCCAAACTTGGAAGTTCCAGA F19 48

>GAL2 (360c6) 1004bp in-house: 625-1004 PathoSeq: 1-624

CTTTGTTCATTTCCGAGGTTTCTCCAAAACATTTGAGAGGTACTTTGGTGTGCTG
TTTCCAATTGATGATTACCTTGGGT
ATCTTCNTGGGNTATTGGCTACCTATGGTACTAAGAGTTACTCAGACTCTAGAC
AATGGAGAATTCCATTAGGTTTATGT
TTCGCCTGGGCTTTATGTTTGGTTGCTGGTATGGTTAGAATGCCAGAATCTCCA
CGTTACCTTGTCGGTAAAGACAGAAT
TGAAGATGCTAAAATGTCACTTGCCAAAACTAACAAGGTTTCTCCAGAGGACC
CAGCATTATACCGTGAACTTCAATTAA
TCCAAGCTGGTGTTGAAAGAAAGATTGGCCGGTAAAGCATCTTGGGGTACT
TTATTCAATGGTAAACCAAGAATCTTT
GAAAGAGTTATTGTTGGTGTCATGTTACAAGCCTTACAACAATT

F19 49 (Con+)

>KGD2 (98c_cp) 334bp in-house: 139-334 public: 1-138

TTCTAACAACAACATCTTTCTTGGATCTTCAATCAATTCCTTGATGGTTCTTAAG
AAAATAACAGCTTCACGACCGTCAA
CTACTCTGTGGTCGTAAGTCAATGCTAAGTACATCATTGGTCTAGAAACGATTT
GTCCGTTAACAGNAATTGGTCTTTNT
TTAAAANTGTGTAAACCAAATACGGNAGTTTAANGCATTTTTATAATTGGGGT
ACAGTATAATGATCCAATAACACNGNC
ATTANAAATAGTGAAAGAACCNCCGGTCATATCTTACAAAGTCAATTTACNAT
TTCTGGCTTTNTTACNCAAATTANANA
TTTCCTTTTNAATA

>MAA (249c af) 619bp

AACCCCACCTTCAAAGACAAAGAAGATTTCGTCAAGCAAACGAATGTCAGAGC AGAAAAGAACCAAGAACTAATCAAATT TGCCCGTGACAACCTTAACCATTTACCATTCACCGAAAAAGACGGAGGTGCAT **GGGAAAACTATGAACGAATGATCAGTG** GTATGCTCTACAACTGTTTACAAAAAGAATTGGAAACAACACGTATGTCTTGC AGAGACTACATGTTGGACTACGGCAGT TTCAGAACTAGAGATTATAAAACAACCCAAGAATTTCTTGATGCAAAATACAA ACATTTAGAAAGTTTCATTGGACATGT TGGCAAAAATGCATTTATGGAATATCCAATCTATTTTGATTATGGGTTTAACAC TTATTTGGGTGATAATTTCTATTCCA ATTACAATTTGACAATTTTGGATGTTTCCATAGTCAGAATTGGTAATAATGTCA AGTGTGGTCCCAATGTATCTATCCTT ACCCCAACACCCCAGTGGATCCCACTTTGCGCTATGATCAATTGGAAAATGC CTTGCCTGTGACGGTGGGTAACGGGGT CTGGTTGTGTGGAAGCTGTACCATTCTTTGGTGGGGTGACAGTANGTGATGGCA GCATT

>MEG1 (55g1) 1380bp in-house: 1-368 public: 497-1096 PathoSeq: 369-496/1097-1380

AATTACAATCTGGTTTGTTACTACCATATCCCATTAGTGTTATTGTCATTGTAGA TATTGATAATGGTTAAAGGATTGGT TTTCATTTTTTGTGTAATGAATGAGCCAAAATAAAAAATCAATTCGATGCGATG CAATGAAGTTTAATAAAATTTTTTT TTTCTTTATTCTTTTAATCAACCCNNCNATCNTTAAATTGAATCAATACCNACC ATTAACATACTTCTATATNCNTATA TTTNTTTTACAAAATATCNTGGGGNAGANAACAACTAGTGNTNCNAAAACAAA ACNACNTCCTTATCCNTTATTAAAANA NATTTCNTCCCCAGGNGGGNATTTAAGAACCGTTCCCAGANNATCATCATCAT CATCATCACAAAAGAAGAAATCATCAA AGAAACAAAGACATAACGATGAAGACGACGAAGAAAATGGTGGCGGTGAAGG ATTTTTAGATGCTTCTAGTTCAAGAAAG ATTTTACAATTGGCAAAAGAACAACAGGATGAATTGGAACAGGAAGATGAAA CACAAAATCACCTTTCATTTGTTCAATC ATTAAAAAATCAACAAATAGATAGTGAAGAAGAAGAAGAAGAAGAAGATGATTAT TCAGATTTGGAAGAAGAAGAAGAAGTTG AAGAGATATTATATGATGAAGAAGATGCAGAATTTATTCCCAAAGATGCAGAA TTATTTAATAAATATTTCCAATCCAGC AGCTGATAAATTTTNAGCCAAAATTCG AGAAANAGAATCCCCACAACAACNACAACAACAGAGTTTTCCAGATAATAGT AATGAAGATGCCGTATTGTTACCACCAA AAGTCATTTTAGCTTATGAAAAAATTGGTCAAATTTTATCAACTTATATTCATG **GGAAATTACCTAAATTATTTAAAATT** TTACCAAGTTTAAAAAATTGGCAAGATGTATTATACGTGACAAATCCAAATAG TTGGACTCCTCATGCCACATATGAAGC AACTAAATTATTTGTGTCGAATTTATCAAGTAATGAAGCTACAGTTTTCATTGA **AACTATCTTGTTGCCACGATTCCGTG** ATTCTATTGAAAATTCCGATGATCATTCATTAAATTATCATATTTATCGAGCATT AAAAAAATCATTATATAAACCAGGA GCTTTTTTCAAAGGGTTCTTGTTACCTTTAGTCGATGGTTATTGTTCTGTACGTG AAGCCACTATTGCTGCTTCAGTGTT AACTAAAGTTTCTGTCCCTGTTTTACATTCATGTCATTATTGTGGCGTACTGATG AATAAAAAACGAGAATCACCTGTAT TTGTCCTACGGCGAATATAA

>RNR1 (38) 2562bp in-house: 1-2562

ATGTATGTTTATAAGAGAGATGGCCGTAAAGAGCCAGTACGTTTCGACAAAAT
CACTGCCAGAGTTCAAAGATTATGTTA
CGGTTTGAATCCAAACCACGTTGAACCAGTTGCTATTACCCAAAAAGTTATATC
AGGTGTTTACCAGGGGGTTACTACTA
TTGAGTTGGACAACTTGGCTGCAGAAATTGCTGCTACAATGACAACAATTCAC
CCAGATTACGCTGTCTTAGCCGCTAGA
ATTGCCGTATCAAATTTACATAAGCAAACCACCAAACAGTATTCCAAAGTGTC
TAAGGATTTATATGAATACATTAATCC
TAAGACTGGGTTACACTCTCCTATGATTTCCAAAGGAAACCTACGACATCATTAT
GGAACACGAAGATGAATTAAACTCAG

CCATTGTTTACGACAGAGATTTTAACTACAATTATTTTGGGTTCAAGACTTTGC
AAAGATCATATTTGTTACGTATCAAC
GGTAAGGTTGCTGAAAGACCACAACATTTGATCATGAGGGTTGCTGTCGGTATTCACGGTAATGATATACCAAGGGTCAT
TGAAACCTATAACTTGATGTCTCAAAGATTCTTCACCCATGGTTCTCCTTGTTT
TITTAACGCTGGTACACCAAGACCAC
AAATGTCCTCATGTTTCTTGCTTGCTATGAAGGATGATTCTATTGAAGGTATTT
ACGACACTTTGAAATCGTGTGCTTTG
ATCTCAAAAAGTGCTGGAGGAATCGGTTTACACATCCACAACATTCGTTCTAC
GGTGCTTACATTGCTGGTACCAATGG
TACTTCTAATGGTATTATTCCAATGGTAAGAGTATTCAATAACACTGCACGTTA
TGTCGACCAAGGTGGTAACAAGAGAC
CTGGTGCCTTTGCCTTGTACTTAGAACCATGGCACAGTGACATTTTTGATTTCA
TTGATATTAGAAAGAATCACGGTAAA
GAAGAAATCAGAGCCAGAGATTTGTTCCCAGCTTTGTGGATTCCAGATTTGTT
ATGAAAAGAGTTGAACAAAATGGTGA
CTGGACTITATTCTCACCAAATGAGGCCCCAGGCTTGGCTGATGTTTATGGTG
CGAATTCGAAGAATTATACACCAAAT
ACGAAAAAGAAAACCGTGGTAGACAGACCATCAAAGCTCAAAAATTGTGGTA
TGCTATTTTGGGAGCCCAAACTGAAACA
GGTACCCCATTTATGTTATATAAAGATTCATGTAACAACAAATCCAACCAA
GAACTTGGGTATTATCAAATCTTCCAA
CTTGTGTTGTGAAATTGTTGAATATTCTGCTCCAGATGAAGTTGCTGTTTGTAA
CTTGGCTTCCATTGCCATCAT
TTGTTGAAAATGATGAAAAAAGTACTTGGTACAACTTTGACAAATTACATCAG
GTCACTAAGGTTGTCACCCGTAACTTG
AACAGAGTTATTGACCGTAACCATTACCCAGTCCCAGAAGCTGAAAGATCAA
CATGAGACACAGACCAATTGCTTTGGG
TGTTCAAGGTTTGGCTGATGCCTTTATGGAATTGAGATTACCATTTGACTCTCA
AGAAGCTAGAGAATTGAACATTCAAA
TTTTTGAGACTATCTACCATGCTGCTGTTGAAGCTTCAATTGAATTGGCTAAAC
AAGAAGGTGCCTACGAAACCTATCCA
GGTTCTCCAGCCTCTCAAGGTTTATTACAATTTGATTTG
ACTGAATTATGGGATTGGGATACATT
AAAACAAGATTTGGCCAAACATGGTATGAGAAACTCCTTGTTGGTTG
TGCCTACTGCTTCCACATCACAAATTT
TGGGTAACAATGAATGTTTTGAACCATACACTTCTAACATTTACTCTAGAAGAG
TATTAGCTGGAGAATTCCAAATTGTC
AATCCATATTTATTGAAGGACTTGGTTGATTTGGGTGTCTGGAACGACGCTATC
AAAAGTAGTATTATTGCTAACAATGG
TTCTATCCAAGCCTTACCAAACATCCCTGATGAAATCAAGGCATTGTACAAAA
CTGTCTGGGAAATCTCACAAAAACATA
TTATCGACATGGCTGCTGATAGAGCAGCATTTATTGATCAATCTCAATCATTAA
ACATTCACATCAAAGATCCAACAATG
GGTAAATTAACCAGTATGCACTTCTACGGTTGGAAGAAAGGTTTAAAGACTGC
TATGTACTACTTAAGAACACAAGCTGC
CAGTGCTGCTATTCAATTTACCATTGATCAAAAGATTGCTGAGACTGCCGGTCA
TA COCTTOCA A A CTTCCA CA A A TTA A
Fig 53 (wn+)
<i>F17</i> - /

ACATTAAGAAATATGTTAACAAAGGAAGAGTTGAGAGTGAGAATACCAGTGAT
GCTCCATACAAGTCACCATCAACCGAA
CCAACCTCATTAGAAAGTTCAGTTGCTGATTTGAAAATAAAAGATGAAGGTGA
AAAGCCAGCTGAAGACAAAACCATTGA
AGAACTCGAAAATGACATTTATAGTGCCAAAGTTATCGCATGTGCTATTGATA
ATCCAGAATCTTGTACAATGTGTTCTG

F19 53 (cont)

>RPL16 (485cL) 759bp in-house: 1-759

GGAGGTNTCNNTCTCTGATTCTTCTCCCCTGCTCCACNCAAGGGCCAACCAACA ATGAGTCAAGTCGCTCCAAAGTGGTA CCAATCAGAAAGACGTTCCAGCTNCAAAACAAACCAGAAAAGACTGCTCGTCC ACAAAATTACGTGCCTCTTTAGTCCC AGGTACCGTTTTAATTTTATTGGCCGGTAGATTCAGAGGTAAAAGAGTTGTTTA CTTGAAGAACTTGGAAGACAACACCT TATTGGTTTCTGGTCCATTCAAAGTCAATGGTGTTCCATTGAGAAGAGTTAACG CTAGATACGTTATCGCCACCTCCACC AAAGTCAACGTTTCTGGTGTTGATGTTTCTAAATTCAACGTCGAATACTTTGCT AGAGAAAAATCTTCTAAATCTAAAAA AAAGAGTTGCTGACCAAAAATCTGTCG ATGCTGCTTTATTAAGTGAAATCAAAAAGACCCCATTATTGAAACAATACTTG GCCGCTTCATTCTCTTTGAAGAACGGT GACAGACCACACTTGTTAAAATTTTAATTTAGGTGAAATTAATATTTTGCAAAC ATGTTCATGATAAATAACAATGNGGG CTTTTTAAAGCAATGGGATGGGGATATGGTTAAGAGGGATGGCTTTATATTTTG AGTTTTTATATATGGGGACCTTTGGT TTAATAAATGGAANGNTATTGGGCTTCAAAATGAACTTN

>RPS21 (328c3) 391bp in-house: 1-391



>RVS167 (67g1) part1 733bp in-house: 145-733 public: 1-144

CTTCAGTTCATATTTAGAAAAATT TCTCNTATGACGATCCAAATCCAATTGTTTATGGTCTCTTTTCACTGACATTTTC CTTATAGCTTGTATAATCTTCAATA ATTCTTGTGCTGGTTCAACAATTCTTTTTCAATCAATTCCAAATCGGGTTTTAA GGTATCTTTGAGATCTTTAACCACT GCTTGGTACAGTTCCGATGCTTCAATACCTTGTGGGTTATCTTCTGGTACCGTA GCACTGGGGTCCGATAATCTACCACT GATTGGTTTATAAATCTCAGCCACGGCTTTGGCAAAATCAATTTGTTCATCTAA CATCCCATTGACAGCATTGAAATATT TCTTGGATTCTTCACTCAACTTTTTTGTTTCCRTTTCGATTTCTTTGAATCTTCTT TCAGCATCGAGATAAACAGCATCT TGGGTGATTTCTCCCATGTTGAATTTCTGACGCATTGTCTGTGGGGCCCTAAGG ACACCCTTTTTGAATCCTTTAAATGA ATAGTATATGAAAAGGAAGGGGCGGAGG GAATTAATTGTAGGAAGAAGTGGCATTGCTTTTTGTCGAAAGCATTTTTTGAGC GTGCGAGAAATTTAATCCAAAAAAAT Fig 56(G) **GTGTGGTGAAAGG**

>RVS167 (67g1) part2 1079bp public: 1-523/668-1079 PathoSeq: 524-667

AGTGGTAGATTATCGGACCCCAGTGCTACGGTACCAGAAGATACCCTACAAGG TATTGAAGCATCGGAACTGTACCAAGC AAGAATTGTTGAACCAGCACAAGAAT TATTGAAGATTATACAAGCTATAAGGAAAATGTCAGTGAAAAGAGACCATAAA CAATTGGATTTGGATCGTCATAAGAGA AATTITTCTAAATATGAACTGAAGAAGAAGAAGAACTGTTAAAGATGAAGAAAA AATGTTCAGTGCTCAAGCAGAAGTAGA AGTTTTGTATCAAATGCAAAGTGATT TTATCAAACCATTGTATGTATCATTCTATTACATGCAGTTGAATATTTTCTACAC ATTATACACTAGAATGGAAGAGTTG AAAATTCCATATTTTATTTTGTCTACTGATATTGTCGATGCNTATACTGCCAAG AAGGGGAACATTGAGGAACAAACCGA TTCTATTGGAATCACTCATTTCAAAGTCGGGCATGCCAAATCCAAATTGGAAGC CACTAAAAGAAGACATGCTGCTATGA AATAGTCCACCTCCTACNGGTGCCAAGCTCTATGGCATCTACAGGAACTGGTG **GTGAATTACCTGCATACTCCCCAGGAG** GTTACAACCAACCATATGGTGATAGCAAGTATCAACCACCATCTTCTCCAGCA ACATACCAATCTCCAGTAGTAGCAGCC ACTGCTCAATCTCCAGCTACTTATCAATCGCCAGTGGCTACTGGACAACCTCCA TCATATTTACCACAAACTCCAGCCAG TGCTCCACCACACAGTTGGTAGTGGCCTTCCAACATGCACGGCTTTATACGA TTATACTGCACAAGCCCAGGGTGACT Fig 56 (5)

••

TGACTTTCCCTGCAGGAGCTGTTATTGAAATTATACAAAGAACCGAAGATGCC AACGGATGGTGGACTGGTAAATACAAT GGTCAAACCGGTGTGTTCCCTGGTAATTATGTGCAATTA F19 56(5) (out)

>SAM2 (36) 1155bp in-house: 1-1155

ATGACTACTTCCAAGGAAACTTTCCTTTTCACTTCAGAATCCGTTGGTGAAGGT CACCCAGATAAGATTTGTGACCAAGT CTCCGATGCCATTTTAGATGCTTGTTTAGCTGTTGATCCATTGTCAAAAGTTGCT TGTGAAACTGCTGCCAAAACCGGTA TGATTATGGTTTTTGGTGAAATTACCACTAAAGCTCAATTGGATTATCAAAAAA TCATTAGAGACACCATTAAACACATT GGTTACGACGATTCTGAAAAAGGTTTTGATTACAAGACTTGTAACGTCTTGGTT **GCAATTGAACAACAATCTCCAGATAT** TGCTCAAGGTTTACATTACGAAAAAGCTTTGGAAGAGTTGGGTGCTGGTGATC AAGGTATTATGTTTGGTTATGCCACCG ATGAAACCGATGAAAAATTGCCATTGACCATTTTATTGGCCCACAAATTGAAT GCTGCCTTGGCTTCTGCCAGAAGATCA GGTTCCTTGCCATGGTTGAGACCAGATACCAAAACCCAAGTCACCATCGAGTA TGAAAAAGATGGTGGTGCAGTTATCCC AAAAAGAGTCGACACAATTGTTATTTCCACTCAACATGCCGAAGAAATCACCA CCGAAAATTTGAGAAAAGAAATTATTG AACATATCATCAAGCAAGTCATCCCAGAACATTTATTAGACGACAAAACTATC **TACCACATTCAGCCATCAGGCAGATTC** GTCATTGGTGGTCCCCAAGGTGATGCTGGTTTGACTGGTAGAAAGATCATTGTT GACACCTATGGTGGTTGGGGTGCACA TGGTGGTGCCTTCTCAGGCAAGGATTTCTCCAAAGTTGATAGGTCTGCTGC TTATGCCGCTCGGTGGGTTGCTAAGT CGTTGGTGACCGCCGGATTGGCCAAAAGGGCCTTGGTGCAGTTCTCCTATGCTA TTGGGGTTGCTGAACCCACCAGCATT TATATAGACACCTATGGGACATCTAAATTGAGCACCGAAGCCCTTGTAGAAAT TATCAAGAATAATTTTGACTTACGCCC TTCTTACGGACATTTTACTAACCAAG Fig 57 AAAATTCTTGGGAACAACCAAAAAAAATTAAAATTT

>SAP (232c_cp) 619bp

AACCTATAATTTTCAGAAAGAGACTAGATTCTGATAGAAATATAGACGCATCA CTATATTTTGGAAATATAGATCCACAA GTTACGGAGTTGTTAATGTATGAGTTGTTCATCCAATTTGGTCCCGTCAAATCA ATCAATATGCCAAAGGATCGTATATT GAAAACACACCAGGGGTATGGATTTGTCGAATTTAAAAAACTCAGCAGATGCCA AATATACTATGGAAATACTACGAGGAA TAAGACTTTATGGAAAAGCATTGAAATTGAAACGAATTGATGCCAAGTCTCAG TCATCAACAAACAACCCAAATAATCAA

>SHA3 (83c3) 1376bp in-house: 375-1376 PathoSeq: 1-374

TGNCCTGGAAATCCCCCATTACCATTTTAAAGGTACCACCACCCCCCAAANCT TNGCGACTATCCATCCAGGTATTANC CCTTGGAGGATTNGCCCATAATAATATGGATGGATCATTTGGAGCAAGGAGAT TTGTCCACTAATATCATGGATAGACAA ATATCCACCAANAATAGTCATAGAAAAGTTCCAAGAACAGATTTTGAANCCCA ATTATTAATGAAGAATGCCATGTTACA ATTGATAGAAGCCATTGAATATTGTCACGAAAATAATATTTACCATTGTGATTT AAAACCAGAAAACATTATGGTTAGAT ATAATCCATACTATGTTCGTCCAACTATCAATAACAATAACAATGGAGAA GATGATTTATGCTATGCCAACAGTATT ATTGACTATAATGAATTACACCTCGTGTTGATTGATTTTGGTTTAGCTATGGAC TCTGCTACCATTTGTTGTAATTCATG TCGTGGATCGTCATTTTACATGGCACCAGAAAGAACCACCAATTATAACACCC **ATCGTTTAATCAACCAATTAATTGATA** TGAATCAATATGAGTCAATTGAAATCAATGGGACAACAGTGACAAAATCAAAC **TGTAAATATTTACCTACATTGGCTGGG** GATATTTGGTCATTGGGAGTATTGTTCATTAATATCACTTGTTCAAGAAACCCA TGGCCCATTGCATCATTTGATAATAA GAGCAAAATCTTACCCATTTCCTCAC AATTTAATCGCTTATTAGATAGAATTTTCAAATTGAATCCTAATGATAGAATAG ATTTACCAACTTTATACAAAGAAGTT **ATTCGTTGTGATTTCTTCAAAGATGATCATTACTACTATGCCCAACATCAACAT** CATCACAATCACAATCAAATCAATAA TGCTTACAATCACTATCAGAAACAACCTAATCAAGCAAGACCTACTGCAAACC AACAATTGTATACACCACCGGAAACCA CCACTTATAATTCATACGCTAGTGATATGGAAGAAGATGAAATTAGTGATGAT GAGTTTTATTCTGATGAAGAAGATGAA GATATTGAAGACTATGAAGAGGAAGAGGAAGAGTATTTTGGTAATGAGCAAC AACAACAACAGCAAGTCACAACAGTGAA TGGTAATTTTGGTCAAGTTAAAGGTACCTGTTATTACGATACCAAAACCAAAA CAACTACATATATAAAACCACCAGCTG CATATACTTTAGAGACGCCTAGTCAAAGTGTTGAATACTGTTAAGTTGTACACA TAAATAATTAATGACAATTAATAATA Fig 59 **ACGATTAATAATATAG**

>TPI1 (233c_cp2) 636bp

AACCAATTTTAGAAACAATGGCTCGTCAATTTTTCGTAGGTGGTAACTTCAAAG CTAACGGTACCAAACAACAAATCACT TCAATCATCGACAACTTGAACAAGGCTGATTTACCAAAGGATGTCGAAGTTGT CATTTGTCCACCCGCCCTTTACCTTGG TTTAGCTGTTGAGCAAAACAAACAACCAACTGTTGCCATTGGTGCTCAAAATG TTTTTGACAAGTCATGTGGTGCTTTCA CTGGTGAAACCTGTGCTTCTCAAATCTTGGATGTTGGTGCCAGCTGGACTTTAA CTGGTCACAGTGAAAGAAGAACCATT ATNAAAGAATCCGATGAATTCATTGCTGAAAAAACCAAGTTTGCCTTGGACAC TGGTGTCAAAGTTATTNTATGTATTGG TGAAACCTTAGAGGAAAGAAAGGTNGTGTCACTTTGGATGTTTGNGCCAGAC AATTGGGATGCTGGTTCCAAGATTGNN TTGATTGGTCAAACATTGNTGGNNCTTACGAACCTGTTTTGGNCAATTGGGTCT GGTTTANCCCGNTNCCCCANAAGATG CTGAAGAACCTACAAGGTNTTAGACTCATTTGGNCAAGANCATTNGTGNCNA ACAACTGAAAAACCNGANTNTNG

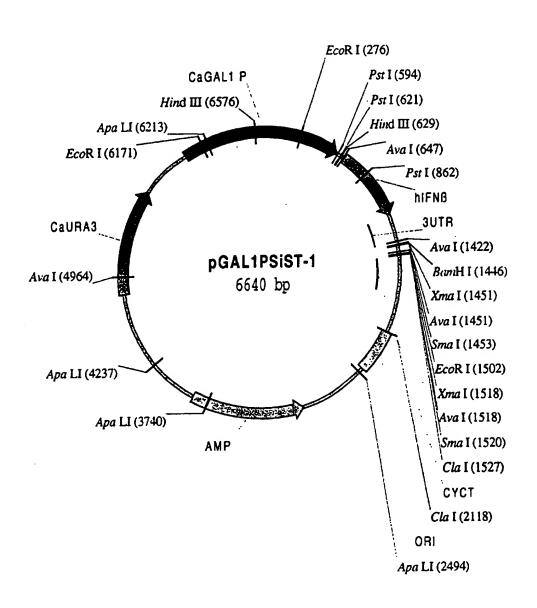


Fig 62

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1	TTCCATCGC AAGGTAGCC	G GAAAGTGGC	GG GGGAAAAAAT CC CCCTTTTTTA	TTTAAGCAC AAATTCGTC	T TCACAAAAC A AGTGTTTTG	C G
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	AAGGTTTTT	T ATATACCTG	CA AAGATGATTG FT TTCTACTAAC	ATAAAAGGG	C TGTGGTTTT	T A
101	CATAATTAA	T TATGAGAAA	G TTAAATGTAA C AATTTACATT	CGTTACAAT	T TATGTTTAT	г
	TGAAGGTGA. ACTTCCACT	A AAGCGATTT. I TTCGCTAAA	A TGATTTTTCC T ACTAAAAAGG	GAAATGAAA CTTTACTTT	A TTTTTTTAC T AAAAAAAT	······································
	GTTTATTTT.	r TTTGTCGGG	C AAAGAAAAAC G TTTCTTTTG	TGAACAAGG	A TTATTAAAA1	· · · · · · · · · · · · · · · · · · ·
			• • • • • • • • • •		. MINNIIII	
			EcoRI			
251	AAAACCACAA	ACAAACACAG	TGGAGAATIC ACCTCTTAAG	TAAGGAGAGA	GTAGAAGTGT	
301	CAATCOTORAC		CGATTCATGA	m. 2000.000	. =	
	GTTACAAATC	TGTAGACTGT	GCTAAGTACT .	ATCAAGCCAA	AGGCCCCAAC	*************************
351	GTGTTTAGTT	TTCGTTTTTC	TTTTTTTTTG	GAAAGAATGT	TTTAGCTCAT	
	ACCAAAAGAA	AGAAGTAAGT	ATAGTTTTGA A	AAGAATTTGC PTCTTAAACG	CCACTTGTTA GGTGAACAAT	•••••••••••
451	TTACAATCAT AATGTTAGTA	ATAAAATTAA TATTTTAATT	ACTTTGATAT A	AAATAGAGT TTTTATCTCA	TTGAAAGTTT AACTTTCAAA	
501	CCCAGATCCT	TTTTGATTTC	TTTGTAAATT T	TTTTTTTCTC	CCACATATAC	••••••
			AAACATTTAA A			
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551 2	TGTGTATGTT	TGGCTAAAAA	ATAAGAAAGA G TATTCTTTCT C	AATATGGGA	CGTCGAGCTG	
		PstI	HindI	11	AvaI	
601 (CTCGACTGTT GAGCTGACAA	TAAACCTGCA	GGCATGCAAG C	TTGGCCAAA	AAGGCCTCGA TTCCGGAGCT	
_	AvaI					
С	CCTTGTACTG (STIGITCACA (CTCCTCCAAA TT GAGGAGGTTT AA	ACGAGAGGA	CAACACGAAG	
						•••••
A	GGTGATGTC C	GAGAAAGGTA C	SAGCTACAAC TI	CGAACCTA A	AGGATGTTTC	
51 A	AGCAGCAAT 1	TTCAGTGTC A	GAAGCTCCT GT CTTCGAGGA CA	GGCAATTG A	ATGGGAGGC	•••••••••••
01 т	TGAATACTG C	CTCAAGGAC A	GGATGAACT TT	GACATCCC 7	GAGGAGATT	***************************************
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851	AAGCAGCTGC	AGCAGTTCCA TCGTCAAGGT	GAAGGAGGAC	GCCGCATIGA	CCATCTATGA	<b>.</b>
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901		AACATCTTTG TTGTAGAAAC				
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		GACTATTGTT				
331		CTGATAACAA				
1.001	ATABACCATC	TGAAGACAGT	CCTGGAAGAA	AAACTGGAGA	AAGAAGATTT	
1001		ACTTCTGTCA				
1051	CACCAGGGGA	AAACTCATGA	GCAGTCTGCA	CCTGÀAAAGA	TATTATGGGA	
		TTTGAGTACT				
1101		TTACCTGAAG				
		AATGGACTTC				
1151		TGGAAATCCT				
		ACCTTTAGGA				
						• • • • • • • • • • • • • • • • • • • •
1201	AGGTTACCTC	CGAAACTGAA	GATCTCCTAG	CCTGTGCCTC	TGGGACTGGA	
		GCTTTGACTT				
1251		AAGCATTCTT TTCGTAAGAA				
		TICGIAAGAA				
		CTGCATATGA				
1301	CCCATTACAT	GACGTATACT	TTCCTGTGAT	CTTCTAAAAC	TTTAAAAATA	
		GTTATTTTTA				
	ATTTAATACT	CAATAAAAAT	TTTAAATAAA	AAAATAAAAC	CTTTTATTTA	
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					SmaI	
					BamHI	
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			AvaI		Aval	
1.401	ma mmmmma-cm	GCAAAAGTCC	 CTCCACCCCT	*CCCCCCCC	TAGAGGATCC	
1401	ATABABACCA	CGTTTTCAGG	GAGCTCCGGA	TCGCCGGCGG	ATCTCCTAGG	
	XmaI					
	~~~~					
	SmaI					
	AvaI					
1451	CCGGGCGCTA	GCCGCCGCT	AGGCCTTTTT	GGCCAAGCTC	GAATTTCGAG	
<b></b>	GGCCCGCGAT	CCGCCGGCGA '	TCCGGAAAAA	CCGGTTCGAG	CTTAAAGCTC	
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		XmaI				
		SmaI	~~			
	EcoRI	AvaI				
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1501	GAATTCGAGC	TCGGTACCCG	GGGATCGAT	CCGTCCCCT	TTTCCTTTGT	
	CTTAAGCTCG	AGCCATGGGC (CCCTAGCTA	GGCAGGGGA	AAAGGAAACA	

1551 (Самамозмо мазама може сель
1551 CGATATCATG TAATTAGTTA TGTCACGCTT ACATTCACGC CCTCCCCCCA GCTATAGTAC ATTAATCAAT ACAGTGCGAA TGTAAGTGCG GGAGGGGGGT
THE ATTACON ACASTICCAN TOTAL STOCK GGAGGGGGGGT
1601 CATCCGCTCT AACCGAAAAG GAAGGAGTTA GACAACCTGA AGTCTAGGTC
GTAGGCGAGA TTGGCTTTTC CTTCCTCAAT CTGTTGGACT TCAGATCCAG

1651 CCTATTTATT TTTTTATAGT TATGTTAGTA TTAAGAACGT TATTTATATT
GGATAAATAA AAAAATATCA ATACAATCAT AATTCTTGCA ATAAATATAA
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1701 TCAAATTTTT CTTTTTTTC TGTACAGACG CGTGTACGCA TGTAACATTA
AGTTTAAAAA GAAAAAAAA ACATGTCTGC GCACATGCGT ACATTGTAAT

1751 TACTGAAAAC CTTGCTTGAG AAGGTTTTGG GACGCTCGAA CCCTTTTAATT
ATGACTITTG GAACGAACTC ITCCAAAACC CTGCGAGCTT CCGAAATTAA
1801 TGCAAGCTAG CTTGGCGTAA TCATGGTCAT AGCTGTTTCC TGTGTGAAAT
ACGITCGATC GAACCGCATT AGTACCAGTA TCGACAAAGG ACACACTITA
1851 TGTTATCCGC TCACAATTCC ACACAACATA CGAGCCGGAA GCATAAAGTG
ACAATAGGCG AGTGTTAAGG TGTGTTGTAT GCTCGGCCTT CGTATTTCAC
1001
1901 TAAAGCCTGG GGTGCCTAAT GAGTGAGCTA ACTCACATTA ATTGCGTTGC ATTTCGGACC CCACGGATTA CTCACTCGAT TGAGTGTAAT TAACGCAACG
TACCCCAACG
1951 GCTCACTGCC CGCTTTCCAG TCGGGAAACC TGTCGTGCCA GAGATCTCTG
CGAGTGACGG GCGAAAGGTC AGCCCTTTGG ACAGCACGGT CTCTAGAGAC
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2001 CATTAATGAA TCGGCCAACG CGCGGGAGA GGCGGTTTGC GTATTGGGCG
GTAATTACTT AGCCGGTTGC GCGCCCCTCT CCGCCAAACG CATAACCCGC

2051 CTCTTCCGCT TCCTCGCTCA CTGACTCGCT GCGCTCGGTC GTTCGCCTCC
GAGAAGGCGA AGGAGCGAGT GACTGAGCGA CGCGAGCCAG CAAGCCGACG
ClaI
2101 GGCGAGCGGT ATCAGATCGA TCTCACTCAA AGGCGGTAAT ACGGTTATCC
CCGCTCGCCA TAGTCTAGCT AGAGTGAGTT TCCGCCATTA TGCCAATAGG
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2151 ACAGAATCAG GGGATAACGC AGGAAAGAAC ATGTGAGCAA AAGGCCAGCA
TGTCTTAGTC CCCTATTGCG TCCTTTCTTG TACACTCGTT TTCCGGTCGT

2201 AAAGGCCAGG AACCGTAAAA AGGCCGCGTT GCTGGCGTTT TTCCATAGGC
TTTCCGGTCC TTGGCATTTT TCCGGCGCAA CGACCGCAAA AAGGTATCCG
2251 TCCGCCCCC TGACGAGCAT CACAAAAATC GACGCTCAAG TCAGAGGTGG
AGGCGGGGG ACTGCTCGTA GTGTTTTTAG CTGCGAGTTC AGTCTCCACC
2301 CCAAACCCCA CACCACCACCACCACCACCACCACCACC
2301 CGAAACCCGA CAGGACTATA AAGATACCAG GCGTTTCCCC CTGGAAGCTC GCTTTGGGCT GTCCTGATAT TTCTATGGTC CGCAAAGGGG GACCTTCGAG
2351 CCTCGTGCGC TCTCCTGTTC CGACCCTGCC GCTTACCGGA TACCTGTCCG
GGAGCACGCG AGAGGACAAG GCTGGGACGG CGAATGGCCT ATGGACAGGC

2401 CCTTTCTCCC TTCGGGAAGC GTGGCGCTTT CTCATAGCTC ACGCTGTAGG
GGAAAGAGGG AAGCCCTTCG CACCGCGAAA GAGTATCGAG TGCGACATCC

ApaLI
2451 TATCTCACTT CCCTCTACCT COTTCCCTCA 22000000
2451 TATCTCAGTT CGGTGTAGGT CGTTCGCTCC AAGCTGGGCT GTGTGCACGA ATAGAGTCAA GCCACATCCA GCAAGCGAGG TTCGACCCGA CACACGTGCT
CACACGIGCT

2501			C GCTGCGCCTT G CGACGCGGAA			
• • •	• • • • • • • •	• • • • • • • •	· · · · · · · · · · ·	• • • • • • • • •		
2551			C GACTTATCGC CTGAATAGCG			
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	TTGTCCTAA	CGTCTCGCT	GTATGTAGGC CATACATCCG	CCACGATGT	C TCAAGAACTT	
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	CACCACCGG	A TTGATGCCG	ACACTAGAAG TGTGATCTTC	CTGTCATAA	CCATAGACGC	
						• • • • • • • • • • • • • • • • • • • •
	GAGACGACTT	CGGTCAATGC	TTCGGAAAAA AAGCCTTTTT	CTCAACCATC	GAGAACTAGG	
	CCGTTTGTTI	GGTGGCGACC	TAGCGGTGGT	AAAAAACAAA	CGTTCGTCGT	
						•
	CTAATGCGCG	TCTTTTTTC	GATCTCAAGA CTAGAGTTCT	TCTAGGAAAC	TAGAAAAGAT	
2851	GCCCAGACT	GCGAGTCACC	AACGAAAACT TTGCTTTTGA	GTGCAATTCC	CTAAAACCAG	
2901	ልጥር ልርልጥዋልጥ	CAAAAAGGAT	CTTCACCTAG	ል ጥርርጥጥጥል ል	בית ב ב ב ב ב ב ב	
			GAAGTGGATC			
2951	ААСТТТТААА	тсаатстааз	GTATATATGA	GTAAACTTGG	TCTGACAGTT	
	TTCAAAATTT	AGTTAGATTT	CATATATACT	CATTTGAACC	AGACTGTCAA	• • • • • • • • • • • • • • • • • • • •
3001	ACCAATGCTT	AATCAGTGAG	GCACCTATCT	CAGCGATCTG	TC-TrATTETT CT-T	•
3402			CGTGGATAGA			
3051	AGTAGGTATC	AACGGACTGA	CCCCGTCGTG GGGGCAGCAC	ATCTATTGAT	GCTATGCCCT	
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	CCCGAATGGT	AGACCGGGGT	GTGCTGCAAT CACGACGTTA	CTATGGCGCT	CTGGGTGCGA	
3151			GCAATAAACC CGTTATTTGG			
3201	CCCACAACTC	COCCOCCAAC	TITTATICCCCC	TO A TO CA COT	CONTRACTOR A CORPOR	
			TTTATCCGCC AAATAGGCGG			
3251			GTAGTTCGCC A			
	TTGTTGCCAT	TGCTACAGGC	ATCGTGGTGT (CACGCTCGTC	GTTTGGTATG	
	GCTTCATTCA	GCTCCGGTTC	CCAACGATCA A	AGGCGAGTTA	CATGATCCCC	
			. 			
3401	CATGTTGTGC	AAAAAAGCGG	TTAGCTCCTT (CGTCCTCCG	ATCGTTGTCA	
	• • • • • • • •			• • • • • • • •		
	CTTCATTCAA	CCGGCGTCAC A	TTATCACTCA 1 AATAGTGAGT #	CCAATACCG	TCGTGACGTA	
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3501 AATTCTCTTA CTGTCATGCC ATCCGTAAGA TGCTTTTCTG TGACTGGTGA TTAAGAGAAT GACAGTACGG TAGGCATTCT ACGAAAAGAC ACTGACCACT
3551 GTACTCAACC AAGTCATTCT GAGAATAGTG TATGCGGCGA CCGAGTTGCT CATGAGTTGG TTCAGTAAGA CTCTTATCAC ATACGCCGCT GGCTCAACGA
3601 CTTGCCCGGC GTCAATACGG GATAATACCG CGCCACATAG CAGAACTTTA GAACGGCCCG CAGTTATGCC CTATTATGGC GCGGTGTATC GTCTTGAAAT
3651 AAAGTGCTCA TCATTGGAAA ACGTTCTTCG GGGCGAAAAC TCTCAAGGAT TTTCACGAGT AGTAACCTTT TGCAAGAAGC CCCGCTTTTG AGAGTTCCTA
ApaLI
3701 CTTACCGCTG TTGAGATCCA GTTCGATGTA ACCCACTCGT GCACCCAACT GAATGGCGAC AACTCTAGGT CAAGCTACAT TGGGTGAGCA CGTGGGTTGA
3751 GATCTTCAGC ATCTTTTACT TTCACCAGCG TTTCTGGGTG AGCAAAAACA CTAGAAGTCG TAGAAAATGA AAGTGGTCGC AAAGACCCAC TCGTTTTTGT
3801 GGAAGGCAAA ATGCCGCAAA AAAGGGAATA AGGGCGACAC GGAAATGTTG CCTTCCGTTT TACGGCGTTT TTTCCCTTAT TCCCGCTGTG CCTTTACAAC
3851 AATACTCATA CTCTTCCTTT TTCAATATTA TTGAAGCATT TATCAGGGTT
TTATGAGTAT GAGAAGGAAA AAGTTATAAT AACTTCGTAA ATAGTCCCAA
3901 ATTGTCTCAT GAGCGGATAC ATATTTGAAT GTATTTAGAA AAATAAACAA TAACAGAGTA CTCGCCTATG TATAAACTTA CATAAATCTT TTTATTTGTT
3951 ATAGGGGTTC CGCGCACATT TCCCCGAAAA GTGCCACCTG ACGTCTAAGA TATCCCCAAG GCGCGTGTAA AGGGGCTTTT CACGGTGGAC TGCAGATTCT
4001 33003777777 77777777
4001 AACCATTATT ATCATGACAT TAACCTATAA AAATAGGCGT ATCACGAGGC TTGGTAATAA TAGTACTGTA ATTGGATATT TTTATCCGCA TAGTGCTCCG
4051 CCTTTCGTCT CGCGCGTTTC GGTGATGACG GTGAAAACCT CTGACACATG GGAAAGCAGA GCGCGCAAAG CCACTACTGC CACTTTTGGA GACTGTGTAC
4101 CAGCTCCCGG AGACGGTCAC AGCTTGTCTG TAAGCGGATG CCGGGAGCAG GTCGAGGGCC TCTGCCAGTG TCGAACAGAC ATTCGCCTAC GGCCCTCGTC
4151 ACAAGCCCGT CAGGGCGCGT CAGCGGGTGT CGGGGCTGGC
TGTTCGGGCA GTCCCCGCACA ACCGCCCACA GCCCCGACCG
ApaLI
4201 TTAACTATGC GGCATCAGAG CAGATTGTAC TGAGAGTGCA CCATATCGAC AATTGATACG CCGTAGTCTC GTCTAACATG ACTCTCACGT GGTATAGCTG
4251 GCTCTCCCTT ATGCGACTCC TGCATTAGGA AGCAGCCCAG TAGTAGGTTG CGAGAGGGAA TACGCTGAGG ACGTAATCCT TCGTCGGGTC ATCATCCAAC
4301 AGGCCGTTGA GCACCGCCGC CGCAAGGAAT GGTGCATGCA AGGAGATGGC TCCGGCAACT CGTGGCGGCG GCGTTCCTTA CCACGTACGT TCCTCTACCG
CONSTRUCT CONSCISES SCRIFTCHIA CCACGIACGI TCCTCTACCG
4351 GCCCAACAGT CCCCCGGCCA CGGGGCCTGC CACCATACCC ACGCCGAAAC CGGGTTGTCA GGGGGCCGGT GCCCCGGACG GTGGTATGGG TGCGGCTTTG
4401 AAGCACTAAT AGGAATTGAT TTGGATGGTA TAAACGGAAA CAAAAAAAAG TTCGTGATTA TCCTTAACTA AACCTACCAT ATTTGCCTTT GTTTTTTTC

4451	AGCTGGTACT TCGACCATGA		TTTTATTAAA A				
		TATAAAACT1	GCATCTAATA	AAACAACTTI	CAACGACATC		
	TGCCATTGAT	TCGTAACACT		TAGTCATTCC	TCTTGTTTGA AGAACAAACT		
		TTTTTGCCGA	TAAAAAAAACG	TTAGAATAAA	CCTGCATATT GGACGTATAA	• • • • • • • • • • • • • • • • • • • •	
	ATACAGATAA TATGTCTATT	CATAATGAAA		TTTTTTTTTTTT	TCTTCAATGA	• • • • • • • • • • • • • • • •	
		GTAAGAAAAT	TTGTAACTAG	TTAAGGACTC	GTTGTTGGGG		
4751		GTTTATATAC CAAATATATG	CGCCCCTTTT GCGGGGAAAA	ACAGTTGAAG TGTCAACTTC	AAAGAAATAG	• • • • • • • • • • • • • •	
		AGCAAACAAA TCGTTTGTTT	AGATATGACA TCTATACTGT	GTCAACACTA CAGTTGTGAT	TCTGGATATC	• • • • • • • • • • • • • • • • • • • •	
4851	TGAGAGAGCA	GAAACTCATG		AGCACAGCGA	TTATTTCGAT		• • • • • • • • • • • • • • • • • • • •
		GAAGAAAACC CTTCTTTTGG	TTAAATACAC	GTAGTTAACT	GCAACTATGG	· · · · · · · · · · · · · · · · · · ·	• • • • • • • • • • • • • • • • • • • •
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	ACTAAGGAGT TGATTCCTCA	AGGAGCTCAA	TTAACTATTT	AATCCAGGAA	TACATACGAA		
5001	AATCAAGACT TTAGTTCTGA	CATATTGATA GTATAACTAT		TTTTTCCTAT	GAATCCACTA		
5051		ATTAGAACTT TAATCTTGAA	AGTGCATTTG	TAGTTAAATA	CTAAAAACTT		
5101		TTGCTGATAT AACGACTATA	TGGTAATACC ACCATTATGG	GTAAAGAAAC CATTTCTTTG	AATATATTGG TTATATAACC		• • • • • • • • • • • • • • • • • • • •
5151	TGGAGTTTAT ACCTCAAATA	AAAATTAGTA TTTTAATCAT	GTTGGGCAGA CAACCCGTCT	TATTACCAAT ATAATGGTTA	GCTCATGGTG CGAGTACCAC	•••••	
5201	TCACTGGGAA AGTGACCCTT	TGGAGTGGTT ACCTCACCAA	GAAGGATTAA CTTCCTAATT	AACAGGGAGC TTGTCCCTCG	TAAAGAAACC ATTTCTTTGG	••••••	
5251	ACCACCAACC .	AAGAGCCAAG TTCTCGGTTC	AGGGTTATTG A	ATGTTAGCTG PACAATCGAC	AATTATCATC TTAATAGTAG	•••••	
5301	AGTGGGATCA TCACCCTAGT	TTAGCATATG	GAGAATATTC (CAAAAAACT AGTTTTTTGA	GTTGAAATTG CAACTTTAAC		
5351	CTAAATCCGA 'GATTTAGGCT '	TAAGGAATTI (ATTCCTTAAA (GTTATTGGAT (TTATTGCCCA .	ACGTGATATG IGCACTATAC		
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5401 GGTGGCCAAG AAGAAGGATT TGATTGGCTT ATTATGACAC CTGGAGTTGG CCACCGGTTC TTCTTCCTAA ACTAACCGAA TAATACTGTG GACCTCAACC
5451 ATTAGATGAT AAAGGTGATG GATTAGGACA ACAATATAGA ACTGTTGATG TAATCTACTA TTTCCACTAC CTAATCCTGT TGTTATATCT TGACAACTAC
5501 AAGTTGTTAG CACTGGAACT GATATTATCA TTGTTGGTAG AGGATTGTTT TTCAACAATC GTGACCTTGA CTATAATAGT AACAACCATC TCCTAACAAA
5551 GGTAAAGGAA GAGATCCAGA TATTGAAGGT AAAAGGTATA GAAATGCTGG CCATTTCCTT CTCTAGGTCT ATAACTTCCA TTTTCCATAT CTTTACGACC

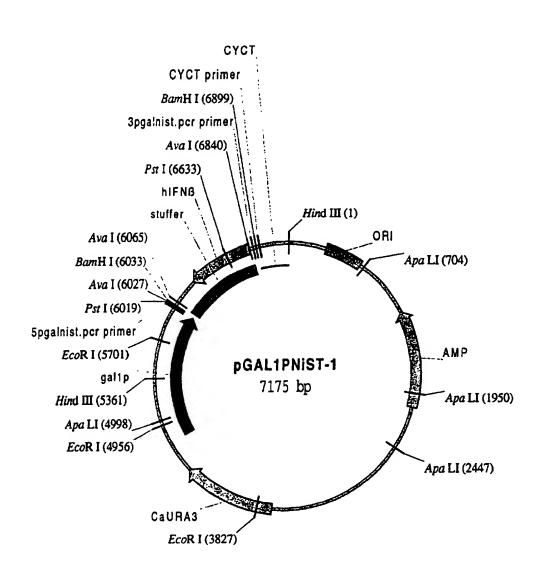
5601 TTGGAATGCT TATTTGAAAA AGACTGGCCA ATTATAAATG TGAAGGGGGA AACCTTACGA ATAAACTTTT TCTGACCGGT TAATATTTAC ACTTCCCCCT
5651 GATTTCACT TTATTAGATT TGTATATATG TAGAATAAAT AAATAAAT
5701 GTTAAATAAA TAATTAAATA AGGGTGGTAA TTATTACTAT TTACAATCAA CAATTTATTT ATTAATTAT TCCCACCATT AATAATGATA AATGTTAGTT
5751 AGGTGGTCCT TCTAGCTGTA ATCCGGGCAG CGCAACGGAA CATTCATCAG TCCACCAGGA AGATCGACAT TAGGCCCGTC GCGTTGCCTT GTAAGTAGTC
5801 TGTAAAAATG GAATCAATAA AGCCCTGCGC TCATGAGCCC GAAGTGGCGA ACATTTTAC CTTAGTTATT TCGGGACGCG AGTACTCGGG CTTCACCGCT

5851 GCCCGATCTT CCCCATCGGT GATGTCGGCG ATATAGGCGC CAGCAACCGC CGGGCTAGAA GGGGTAGCCA CTACAGCCGC TATATCCGCG GTCGTTGGCG
5901 ACCTGTGGCG CCGCAGCGCG CAGGGTCAGC CTGAATACGC GTTTAATGAC TGGACACCGC GGCGTCGCGC GTCCCAGTCG GACTTATGCG CAAATTACTG
5951 CAGCACAGTC GTGATGGCAA GGTCAGAATA GCCCAAGTCG GCCGAGGGGC GTCGTGTCAG CACTACCGTT CCAGTCTTAT CGGGTTCAGC CGGCTCCCCG

6001 CTGTACAGTG AGGGAAGATC TGATATTGAC GAAGAGGAAC CAATGTAACG GACATGTCAC TCCCTTCTAG ACTATAACTG CTTCTCCTTG GTTACATTGC
6051 TTACACTGAA GAAAACACAC AATAAACGGG AAGAAACGGT GTAAAAGTGT AATGTGACTT CTTTTGTGTG TTATTTGCCC TTCTTTGCCA CATTTTCACA

6101 GAAAATAATT TTTGAATATC ATTTCCCTTG GTTTAATTCC AAACGAAACG
EcoRI
6151 TGTTTTTTT AGAGAATGGG AATTCTTATT GGATGTCTAG ATTGTTTGTT ACAAAAAAA TCTCTTACCC TTAAGAATAA CCTACAGATC TAACAAACAA
ApaLI
6201 TACTCCAGAC TGTGCACAAA AACGTTTGGA TGGATGATCA GAAGATATTT
ATGAGGTCTG ACACGTGTTT TTGCAAACCT ACCTACTAGT CTTCTATAAA
6251 TTAGGCTTAG CTCTAAATAT AAGAAATGAT GCTTGAAAAA CCAGACAGAA AATCCGAATC GAGATTTATA TTCTTTACTA CGAACTTTTT GGTCTGTCTT
6301 ATTGAGTTTC AAAAATTGGT AATGTGAGGT ATTAGTCAAC TAACCAAATA TAACTCAAAG TTTTTAACCA TTACACTCCA TAATCAGTTG ATTGGTTTAT

6351			CATTTCATTT		
	TGTTACGTTT	GGCCAACTAT	GTAAAGTAAA		CTPTGACCTT
6401	TTGGATGACC	AGCACACAAA	CACATAAAGT	AATTATGGGA	ATTAGAAGCG
			GTGTATTTCA		
	,5,00,,,,				
6451	AACATAGAGG				
	TTGTATCTCC	TCATGAACCG	GTGCTTGTCT	TATGTTCACC	CTTGTGATAA
6501	TTCTCCATTG				
	AAGAGGTAAC	AAAATCAAGA	CAAAAAAAACA	GTCGGATCAA	AACACGATAC
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			HindIII		
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6551	TGTAAAAAAT	ATTGCCAAGA	AAAAAAGCTT	GTTTTGTGGC	CAGTGTCCGA
	ACATTTTTTA	TAACGGTTCT	TTTTTTCGAA	CAAAACACCG	GTCACAGGCT
	· · · · · · · · ·				
6601	AAAAAATTTT				
	TTTTTTAAAA	CCCCTTAGAA	GCCTAATTAA	ATACAAAAGT	
		<b></b> .			



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HindIII				
TCGAACTCAT A	TTCTATAGTG TCACCTAAAT AAGATATCAC AGTGGATTTA	TCGAACCGCA	AATCATGGTC TTAGTACCAG	
51 ATAGCTGTTT C	CCTGTGTGAA ATTGTTATCC GGACACACTT TAACAATAGG	GCTCACAATT CGAGTGTTAA	GGTGTGTTGT	
101 TACGAGCCGG A ATGCTCGGCC T	AAGCATAAAG TGTAAAGCCT TTCGTATTTC ACATTTCGGA	GGGGTGCCTA CCCCACGGAT	ATGAGTGAGC TACTCACTCG	
151 TAACTCACAT T ATTGAGTGTA A	PAATTGCGTT GCGCTCACTG ATTAACGCAA CGCGAGTGAC	CCCGCTTTCC GGGCGAAAGG	AGTCGGGAAA	
201 CCTGTCGTGC C GGACAGCACG G	CAGCTGCATT AATGAATCGG	CCAACGCGCG		
251 GTTTGCGTAT T CAAACGCATA A	GGGCGCTCT TCCGCTTCCT CCCGCGAGA AGGCGAAGGA			
301 TCGGTCGTTC G	GCTGCGGCG AGCGGTATCA CGACGCCGC TCGCCATAGT			
TGCCAATAGG T	CAGAATCAG GGGATAACGC GTCTTAGTC CCCTATTGCG	TCCTTTCTTG		
401 AAGGCCAGCA A TTCCGGTCGT T	AAGGCCAGG AACCGTAAAA TTCCGGTCC TTGGCATTTT	AGGCCGCGTT TCCGGCGCAA	CGACCGCAAA	•••••••••••••••••••••••••••••••••••••••
451 TTCCATAGGC TO AAGGTATCCG AG	CCGCCCCC TGACGAGCAT GGCGGGGG ACTGCTCGTA	CACAAAAATC GTGTTTTTAG	GACGCTCAAG	:
501 TCAGAGGTGG CO	GAAACCCGA CAGGACTATA CTTTGGGCT GTCCTGATAT	AAGATACCAG		······································
GACCTTCGAG GO	CTCGTGCGC TCTCCTGTTC GAGCACGCG AGAGGACAAG	GCTGGGACGG		
601 TACCTGTCCG CO	CTTTCTCCC TTCGGGAAGC GAAAGAGGG AAGCCCTTCG	GTGGCGCTTT		ý.
651 ACGCTGTAGG TA	ATCTCAGTT CGGTGTAGGT FAGAGTCAA GCCACATCCA	GCAAGCGAGG	TTCGACCCGA	
ApaLI		· • • • • • • • • • • • • • • • • • • •	• • • • • • • • •	
CACACGTGCT TG	CCCCCGTT CAGCCCGACC (	CGACGCGGAA	TAGGCCATTG	
751 TATCGTCTTG AG ATAGCAGAAC TC	ETCCAACCC GGTAAGACAC ( AGGTTGGG CCATTCTGTG (	GACTTATCGC ( CTGAATAGCG (	CACTGGCAGC GTGACCGTCG	
801 AGCCACTGGT AA TCGGTGACCA TT	CAGGATTA GCAGAGCGAG (	GTATGTAGGC ( CATACATCCG (	GGTGCTACAG CCACGATGTC	
851 AGTTCTTGAA GT TCAAGAACTT CA	GGTGGCCT AACTACGGCT A	ACACTAGAAG ( IGTGATCTTC (	GACAGTATTT CTGTCATAAA	
901 GGTATCTGCG CT CCATAGACGC GA	CTGCTGAA GCCAGTTACC 1 GACGACTT CGGTCAATGG 1	TTCGGAAAAA ( AAGCCTTTTT (	GAGTTGGTAG CTCAACCATC	

951 CTCTTGATCC GGCAAACAAA CCACCGCTGG TAGCGGTGGT TTTTTTGTTT GAGAACTAGG CCGTTTGTTT GGTGGCGACC ATCGCCACCA AAAAAACAAA
1001 - CCARCOACOA CARRACCOCCO AGANANANA CARRACCAR A CAR
1001 GCAAGCAGCA GATTACGCGC AGAAAAAAAG GATCTCAAGA AGATCCTTTG CGTTCGTCGT CTAATGCGCG TCTTTTTTC CTAGAGTTCT TCTAGGAAAC
1051 ATCTTTCTA CGGGGTCTGA CGCTCAGTGG AACGAAAACT CACGTTAAGG TAGAAAAGAT GCCCCAGACT GCGAGTCACC TTGCTTTTGA GTGCAATTCC
1101 GATTTTGGTC ATGAGATTAT CAAAAAGGAT CTTCACCTAG ATCCTTTTAA
CTAAAACCAG TACTCTAATA GTTTTTCCTA GAAGTGGATC TAGGAAAATT
4464
1151 ATTAAAAATG AAGTTTTAAA TCAATCTAAA GTATATATGA GTAAACTTGG TAATTTTTAC TTCAAAATTT AGTTAGATTT CATATATACT CATTTGAACC
1201 TCTGACAGTT ACCAATGCTT AATCAGTGAG GCACCTATCT CAGCGATCTG AGACTGTCAA TGGTTACGAA TTAGTCACTC CGTGGATAGA GTCGCTAGAC
1251 TCTATTTCGT TCATCCATAG TTGCCTGACT CCCCGTCGTG TAGATAACTA AGATAAAGCA AGTAGGTATC AACGGACTGA GGGGCAGCAC ATCTATTGAT
1301 CGATACGGA GGCTTACCA TCTGGCCCCA GTGCTGCAAT GATACCGCGA
GCTATGCCCT CCCGAATGGT AGACCGGGGT CACGACGTTA CTATGGCGCT
1351 GACCCACGCT CACCGGCTCC AGATTTATCA GCAATAAACC AGCCAGCCGG
CTGGGTGCGA GTGGCCGAGG TCTAAATAGT CGTTATTTGG TCGGTCGGCC
1401 NAGGOOGIA GGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG
1401 AAGGGCCGAG CGCAGAAGTG GTCCTGCAAC TTTATCCGCC TCCATCCAGT TTCCCGGCTC GCGTCTTCAC CAGGACGTTG AAATAGGCGG AGGTAGGTCA
1451 CTATTAATTG TTGCCGGGAA GCTAGAGTAA GTAGTTCGCC AGTTAATAGT GATAATTAAC AACGGCCCTT CGATCTCATT CATCAAGCGG TCAATTATCA
1501 TTGCGCAACG TTGTTGCCAT TGCTACAGGC ATCGTGGTGT CACGCTCGTC
AACGCGTTGC AACAACGGTA ACGATGTCCG TAGCACCACA GTGCGAGCAG
1551 GTTTGGTATG GCTTCATTCA GCTCCGGTTC CCAACGATCA AGGCGAGTTA
CAAACCATAC CGAAGTAAGT CGAGGCCAAG GGTTGCTAGT TCCGCTCAAT
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GTACTAGGGG GTACAACACG TTTTTCGCC AATCGAGGAA GCCAGGAGGC
1651 1800889990 011081 088 089000000 080000000000
1651 ATCGTTGTCA GAAGTAAGTT GGCCGCAGTG TTATCACTCA TGGTTATGGC TAGCAACAGT CTTCATTCAA CCGGCGTCAC AATAGTGAGT ACCAATACCG
1701 1702 (2007)
1701 AGCACTGCAT AATTCTCTTA CTGTCATGCC ATCCGTAAGA TGCTTTTCTG TCGTGACGTA TTAAGAGAAT GACAGTACGG TAGGCATTCT ACGAAAAGAC
1751 TGACTGGTGA GTACTCAACC AAGTCATTCT GAGAATAGTG TATGCGGCGA
ACTGACCACT CATGAGTTGG TTCAGTAAGA CTCTTATCAC ATACGCCGCT
1901 CCCACCORCCO COMCCCCCC COCCAMANAGO CAMANAGO COCCAMANAGO COCCAMANAGO CAMANAGO CAM
1801 CCGAGTTGCT CTTGCCCGGC GTCAATACGG GATAATACCG CGCCACATAG GGCTCAACGA GAACGGCCG CAGTTATGCC CTATTATGGC GCGGTGTATC
1951 CACAACTETTA AAACTECCTCA TCATTCCAAA ACCTECTTCA CCCCCCCAACTETTCA
1851 CAGAACTITA AAAGTGCTCA TCATTGGAAA ACGTTCTTCG GGGCGAAAAC GTCTTGAAAT TTTCACGAGT AGTAACCTTT TGCAAGAAGC CCCGCTTTTG

					ApaLI	
	TCTCAAGGAT AGAGTTCCTA	GAATGGCGAC	AACTCTAGGT	CAAGCTACAT	TGGGTGAGCA	
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-	GCACCCAACT CGTGGGTTGA	CTAGAAGTCG	TAGAAAATGA	AAGTGGTCGC	AAAGACCCAC	
	AGCAAAAACA	GGAAGGCAAA	ATGCCGCAAA		AGGGCGACAC	
	GGAAATGTTG CCTTTACAAC	TTATGAGTAT	GAGAAGGAAA	AAGTTATAAT	AACTTCGTAA	•
	TATCAGGGTT					
	ATAGTCCCAA	TAACAGAGTA	CTCGCCTATG	TATAAACTTA	CATAAATCIT	
2151	AAATAAACAA TTTATTTGTT	ATAGGGGTTC TATCCCCAAG	CGCGCACATT GCGCGTGTAA	TCCCCGAAAA AGGGGCTTTT	CACGGTGGAC	
	ACGTCTAAGA	AACCATTATT	ATCATGACAT		AAATAGGCGT	
2251	ATCACGAGGC			• • • • • • • •		
2251	TAGTGCTCCG	GGAAAGCAGA	GCGCGCAAAG	CCACTACTGC	CACTTTTGGA	
	CTGACACATG GACTGTGTAC	CAGCTCCCGG GTCGAGGGCC	AGACGGTCAC TCTGCCAGTG		TAAGCGGATG	
	CCGGGAGCAG GGCCCTCGTC	ACAAGCCCGT	CAGGGCGCGT	CAGCGGGTGT GTCGCCCACA	ACCGCCCACA	
				• • • • • • • •	ApaLI	
2401	CGGGGCTGGC GCCCGACCG	TTAACTATGC AATTGATACG	GGCATCAGAG CCGTAGTCTC	CAGATTGTAC GTCTAACATG	TGAGAGTGCA ACTCTCACGT	
	ApaLI					
2451	CCATATGCGG GGTATACGCC	TGTGAAATAC ACACTTTATG	CGCACAGATG GCGTGTCTAC	CGTAAGGAGA GCATTCCTCT	AAATACCGCA TTTATGGCGT	
2501	TCAGGCGAAA AGTCCGCTTT	AACATTTGCA	ATTATAAAAC	AATTTTAAGC	GCAATTTATA	
 2551	TTGTTAAATC	AGCTCATTTT	TTAACCAATA	• • • • • • • •	GGCAAAATCC	
					· · · · · · · · ·	
	CTTATAAATC GAATATTTAG	TTTTCTTATC	TGGCTCTATC	CCAACTCACA	ACAAGGTCAA	
2651	TGGAACAAGA ACCTTGTTCT	GTCCACTATT CAGGTGATAA	AAAGAACGTG TTTCTTGCAC	GACTCCAACG CTGAGGTTGC	TCAAAGGGCG AGTTTCCCGC	
2701	AAAAACCGTC TTTTTGGCAG	TATCAGGGCG ATAGTCCCGC	ATGGCCCACT TACCGGGTGA	ACGTGAACCA TGCACTTGGT	TCACCCAAAT AGTGGGTTTA	
	CAAGTTTTTT GTTCAAAAAA	GCGGTCGAGG	TGCCGTAAAG	CTCTAAATCG	GAACCCTAAA	

		C GATTTAGAGO				
		AAGAAAGCGA TTCTTTCGCT				
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2	TCGCCAGTC	GCTGCGCGTA GCGACGCGCAT	TGGTGGTGTG	GGCGGCGCGA	ATTACGCGGC	:
						• • • • • • • • • • • • • • • • • • • •
G	SATGTCCCGC	GCAGGTAAGC	GGTAAGTCCG	ACGCGTTGAC	AACCCTTCCC	:
						· · · · · · · · · · · · · · · · · · ·
G	CTAGCCACG	GGGCCTCTTC CCCGGAGAAG	CGATAATGCG	GTCGACCGCT	TTCCCCCTAC	
		CGATTAAGTT				·
A	CGACGTTCC	GCTAATTCAA	CCCATTGCGG	TCCCAAAAGG	GTCAGTGCTG	
3101 G	ייים אם אביייייייייייייייייייייייייייייי	GACGGCCAGT	GAATTGTAAT	ACGACTCACT	<b>АТАСССССАА</b>	
		CTGCCGGTCA				
		. <b></b>				• • • • • • • • • • • • • • • • • • • •
		AATGATGAGC TTACTACTCG				
• • • • •			• • • • • • • • •	• • • • • • • •	• • • • • • • •	•••••
A	ATAGGGCAC	TTGACGCCGG AACTGCGGCC	CGTTCTCGTT	GAGCCAGCGG	CGTATGTGAT	
						***************************************
A	AGAGTCTTA	GACTTGGTTG	TCATGATTAT	CCTTAACTAA	ACCTACCATA	
						• • • • • • • • • • • • • • • • • • • •
T	TTGCCTTTG	AAAAAAAAGA	CGACCATGAT	GAAAGAAATT	TTAATAAAAT	
A	ATAAACTAA	TTATTTAATA AATAAATTAT	CATATATAAT	ATAAAACTTG	CATCTAATAA	
		TTGCTGTAGT				
	<del></del>	AACGACATCA				
			. <b></b> .			• • • • • • • • • • • • • • • • • • • •
		CTTGTTTGAT GAACAAACTA				
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		CTGCATATTA C GACGTATAAT				
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AA	AAAAACAA	CTTCAATGAT ( GAAGTTACTA (	TAAAGTTGG '	TAAGAAAATT	TGTAACTAGT	
						• • • • • • • • • • • • • • • • • • • •
TA	AGGACTCG	AACAACCCCA TTGTTGGGGT /	ATGTGTGACC	AAATATATGG	CGGGGAAAAT	
GT	CAACTTCT	AAGAAATAGA / TTCTTTATCT 1	TATCTTTAT (	CGTTTGTTTT	CTATACTGTC	
		GACCTATAGT C				
		CTGGATATCA C				
	• • • • • • •				• • • • • • • •	
3751 GC	ACAGCGAT '	TATTTCGATT A	ATGGAACTG A	AGAAAACCA	ATTTATGTGC	
CG	TGTCGCTA A	ataaagctaa t	TACCTTGAC 1	TCTTTTGGT	TAAATACACG	

#### EcoRI

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		CAACTATGG	r gattccttaa	GGAACTTAAT	TAACTATTTA	
3851	TAGGTCCTTA ATCCAGGAAT		TAGTTCTGAG			
						• • • • • • • • • • • • • • • • • • • •
		TTAGGTGAT	ACTTGGTAAT	AATCTTGAAA	GTGCATTTGT	
						• • • • • • • • • • • • • • • • • • • •
3951	TCAATTTATG AGTTAAATAC		ATAGAAAATT TATCTTTTAA			
						• • • • • • • • • • • • • • • • • • • •
		TATATAACC	CCTCAAATAT	TTTAATCATC	AACCCGTCTA	
4051	ATTACCAATG TAATGGTTAC		CACTGGGAAT GTGACCCTTA			• .
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4101	ACAGGGAGCT TGTCCCTCGA		CCACCAACCA			
4151	TGTTAGCTGA		GTGGGATCAT CACCCTAGTA			
4201	CAAAAAACTG		TAAATCCGAT			
		GCACTATACC	CACCGGTTCT	TCTTCCTAAA	CTAACCGAAT	
		• • • • • • • • •	• • • • • • • •		• • • • • • • •	• • • • • • • • • • • • • • • • • • • •
4301	TTATGACACC AATACTGTGG		TTAGATGATA AATCTACTAT			
• • • •	· · · · · · · · · ·	• • • • • • • • •	• • • • • • • •			• • • • • • • • • • • • • • • • • • • •
4351	CAATATAGAA GTTATATCTT		AGTTGTTAGC TCAACAATCG		TATAATAGTA	· · · · · · · · · · · · · · · · · · ·
• • • •	• • • • • • • • •		• • • • • • • •			• • • • • • • • • • • • • • • • • • • •
4401	TGTTGGTAGA		GTAAAGGAAG CATTTCCTTC			
4451	AAAGGTATAG					
	FITCCATATC		ACCTTACGAA	INDUCTITIT		
4501	TTATAAATGT AATATTTACA		ATTTTCACTT TAAAAGTGAA			
		TTATTTATTC	AATTTATTTA	TTAATTTATT	CCCACCATTA	
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		ATGTTAGTTT	CCACCAGGAA	GATCGACATT	AGGCCCGTCG	
		TAAGTAGTCA	CATTTTTACC	TTAGTTATTT	CGGGACGCGT	
		AGTCGGACTT	ATGCGCAAAT '	TACTGGTCGT	GTCAGCACTA	
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4751 GGCAAGGTCA GAATAGCCCA AGTCGGCCGA GGGGCCTGTA CAGTGAGGGA CCGTTCCAGT CTTATCGGGT TCAGCCGGCT CCCCGGACAT GTCACTCCCT
4801 AGATCTGATA TTGACGAAGA GGAACCAATG TAACGTTACA CTGAAGAAAA TCTAGACTAT AACTGCTTCT CCTTGGTTAC ATTGCAATGT GACTTCTTTT
•••••••••••••
4851 CACACAATAA ACGGGAAGAA ACGGTGTAAA AGTGTGAAAA TAATTTTTGA GTGTGTTATT TGCCCCTTCTT TGCCACATTT TCACACTTTT ATTAAAAACT
4901 ATATCATTTC CCTTGGTTTA ATTCCAAACG AAACGTGTTT TTTTTAGAGA TATAGTAAAG GGAACCAAAT TAAGGTTTGC TTTGCACAAA AAAAATCTCT
•••••••••••••••••••••••••••••••••••••••
EcoRI ApaLI
4951 ATGGGAATTC TTATTGGATG TCTAGATTGT TTGTTTACTC CAGACTGTGC TACCCTTAAG AATAACCTAC AGATCTAACA AACAAATGAG GTCTGACACG
ApaLI
5001 ACAAAAACGT TTGGATGGAT GATCAGAAGA TATTTTTAGG CTTAGCTCTA TGTTTTTGCA AACCTACCTA CTAGTCTTCT ATAAAAATCC GAATCGAGAT
5051 AATATAAGAA ATGATGCTTG AAAAACCAGA CAGAAATTGA GTTTCAAAAA TTATATTCTT TACTACGAAC TTTTTGGTCT GTCTTTAACT CAAAGTTTTT

5101 TTGGTAATGT GAGGTATTAG TCAACTAACC AAATAACAAT GCAAACCGGT AACCATTACA CTCCATAATC AGTTGATTGG TTTATTGTTA CGTTTGGCCA
5151 mcamacamm cammorcasa amangasa mcaaampen marena
5151 TGATACATTT CATTTTGAAA ATAATGAAAC TGGAATTGGA TGACCAGCAC ACTATGTAAA GTAAAACTTT TATTACTTTG ACCTTAACCT ACTGGTCGTG
5201 ACAAACACAT AAAGTAATTA TGGGAATTAG AAGCGAACAT AGAGGAGTAC TGTTTGTGTA TTTCATTAAT ACCCTTAATC TTCGCTTGTA TCTCCTCATG
5251 TTGGCCACGA ACAGAATACA AGTGGGAACA CTATTTTCTC CATTGTTTTA AACCGGTGCT TGTCTTATGT TCACCCTTGT GATAAAAGAG GTAACAAAAT
5301 GTTCTGTTTT TTTGTCAGCC TAGTTTTGTG CTATGTGTAA AAAATATTGC CAAGACAAAA AAACAGTCGG ATCAAAACAC GATACACATT TTTTATAACG
HindIII
niidii
5351 CAAGAAAAA AGCTTGTTTT GTGGCCAGTG TCCGAAAAAA ATTTTGGGGA GTTCTTTTTT TCGAACAAAA CACCGGTCAC AGGCTTTTTT TAAAACCCCT
F.103
5401 ATCTTCGGAT TAATTTATGT TTTCATTCCA TCGGGGAAAG TGGGGGGGAA TAGAAGCCTA ATTAAATACA AAAGTAAGGT AGCCCCTTTC ACCCCCCCTT
5451 AAAATTTTAA GCAGTTCACA AAACCTTCCA AAAAATATAT GGACAAAGAT
TTTTAAAATT CGTCAAGTGT TTTGGAAGGT TTTTTATATA CCTGTTTCTA
5501 GATTGTATTT TCCCGACACC AAAATCATAA TTAATTATGA GAAAGTTAAA
CTAACATAAA AGGGCTGTGG TTTTAGTATT AATTAATACT CTTTCAATTT
5551 TGTAACGTTA CAATTTATGT TTATTTGAAG GTGAAAAGCG ATTTATGATT
ACATTGCAAT GTTAAATACA AATAAACTTC CACTTTTCGC TAAATACTAA
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5601 TTTCCGAAAT GAAAATTTTT TTTAGGTTTA TTTTTTTTGT CGGGCAAAGA AAAGGCTTTA CTTTTAAAAA AAATCCAAAT AAAAAAAACA GCCCGTTTCT

					EcoRI	
		TTCCTAATAA	TTTTAAAAAC			
• • •	EcoRI		· · · · · · · · · · ·	• • • • • • • •	· • • • • • • • •	
5701		TCTCTCATCT	AGTGTGTTAC	AAATCTGTAG	ACTGTGCTAA	
5751	CATGATAGTT		GGTTGGTGTT	TAGTTTTCGT	TTTTCTTTTT	
5801	TTTTGGAAAG AAAACCTTTC	AATGTTTTAG TTACAAAATC				
5851	TTTGAAAGAA	TTTGCCCACT AAACGGGTGA				······································
5901	GATATAAAAT		• • • • • • • •		• • • • • • • •	:
		TCTCAAACTT				
5951	AAATTTTTTT TTTAAAAAAA	TTCTCCCACA AAGAGGGTGT				
- • •		PstI	AvaI	BamHI		•••••••••••
6001	AAAGAGTTAT TTTCTCAATA	ACCCTGCAGC TGGGACGTCG				
		AvaI				
6051	GCGGCCGCTA CGCCGGCGAT	GGCCTCGAGG CCGGAGCTCC				
	AAAATAAAAT					
9101		AATTTATTTA				
6151	AAAATCTTCT TTTTAGAAGA	AGTGTCCTTT TCACAGGAAA				
• • • •						• • • • • • • • • • • • • • • • • • • •
6201	AAACAGCATC TTTGTCGTAG	ACGACCAACT				
6251	AGGCACAGGC TCCGTGTCCG	TAGGAGATCT ATCCTCTAGA				
6301	AATGAAGTAA TTACTTCATT					
						• • • • • • • • • • • • • • • • • • • •
	AGTGACTGTA TCACTGACAT	GAGGAACCGG	AAGTCCATTA	CGTCTTAGGA	GGGTATTATA	
6401	CTTTTCAGGT GAAAAGTCCA	GCAGACTGCT (CGTCTGACGA (CATGAGTTTT (GTACTCAAAA (CCCCTGGTGA GGGGACCACT	AATCTTCTTT TTAGAAGAAA	•
6451	CTCCAGTTTT GAGGTCAAAA	TCTTCCAGGA (TGTCTTCAG	ATGGTTTATC FACCAAATAG	TGATGATAGA ACTACTATCT	
	• • • • • • • •		• • • • • • • • •		· • • • • • • • • • • • • • • • • • • •	
	CATTAGCCAG GTAATCGGTC	CTCCAAGAGT	GTTATCAGA (GTAAGGTCGG	TCACGATCTA	
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	GAATCTTGTC TGAAAATAGC AAAGATGTTC TGGAGCATCT CATAGATGGT CTTAGAACAG ACTTTTATCG TTTCTACAAG ACCTCGTAGA GTATCTACCA
	PstI
	CAATGCGGCG TCCTCCTTCT GGAACTGCTG CAGCTGCTTA ATCTCCTCAG GTTACGCCGC AGGAGGAAGA CCTTGACGAC GTCGACGAAT TAGAGGAGTC
	GGATGTCAAA GTTCATCCTG TCCTTGAGGC AGTATTCAAG CCTCCCATTC CCTACAGTTT CAAGTAGGAC AGGAACTCCG TCATAAGTTC GGAGGGTAAG
6701	AATTGCCACA GGAGCTTCTG ACACTGAAAA TTGCTGCTTC TTTGTAGGAA TTAACGGTGT CCTCGAAGAC TGTGACTTTT AACGACGAAG AAACATCCTT
• • •	
6751	TCCAAGCAAG TTGTAGCTCA TGGAAAGAGC TGTAGTGGAG AAGCACAACA AGGTTCGTTC AACATCGAGT ACCTTTCTCG ACATCACCTC TTCGTGTTGT

	AvaI
6801	GGAGAGCAAT TTGGAGGAGA CACTTGTTGG TCATGTTCCT CGAGGCCTTT CCTCTCGTTA AACCTCCTCT GTGAACAACC AGTACAAGGA GCTCCGGAAA
• • •	
	BamHI
6951	TTGGCCAGCT GGCGCCTGCT GCGCGACGGC GAGCTGCTCA CCACCCAGGA
	AACCGGTCGA CCGCGGACGA CGCGCTGCCG CTCGACGAGT GGTGGGTCCT
• • • •	······································
	BamHI
6901	TCCGTCCCCC TTTTCCTTTG TCGATATCAT GTAATTAGTT ATGTCACGCT
	AGGCAGGGGG AAAAGGAAAC AGCTATAGTA CATTAATCAA TACAGTGCGA
6951	TACATTCACG CCCTCCCCC ACATCCGCTC TAACCGAAAA GGAAGGAGTT
	ATGTAAGTGC GGGAGGGGG TGTAGGCGAG ATTGGCTTTT CCTTCCTCAA
• • •	
7001	AGACAACCTG AAGTCTAGGT CCCTATTTAT TTTTTTATAG TTATGTTAGT
	TCTGTTGGAC TTCAGATCCA GGGATAAATA AAAAAATATC AATACAATCA
7051	ATTAAGAACG TTATTTATAT TTCAAATTTT TCTTTTTTTT CTGTACAGAC TAATTCTTGC AATAAATATA AAGTTTAAAA AGAAAAAAAA GACATGTCTG
• • • •	
7101	GCGTGTACGC ATGTAACATT ATACTGAAAA CCTTGCTTGA GAAGGTTTTG CGCACATGCG TACATTGTAA TATGACTTTT GGAACGAACT CTTCCAAAAC
• • • •	
	HindIII
7151	GGACGCTCGA AGGCTTTAAT ITGCA
	CCTGCGAGCT TCCGAAATTA AACGT
• • • •	

701 ANSGWXG

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1	MSITVTFPK	s PSTKKRAPA	= = F GIELEFSQQG		LAYPYFSVDN
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51	_	A KYWGYPSSY(LIVKLVKCAN	IEKSQILKTD	_
101	DLIBEADTK	CLFYISLPL(/ YSRIENKKVF	YVLREPEQPK	VSKAPTCERP
151	ASVVAAEEDI	O ENLOGGEEDE	DICEGMOSCV 3	nsgelskgyk	HWHKDHPKYI
201	NDDRVTIGQV	FHQYGLDPS1	PLTHSLFNSI	nemektnääk	nfgvsgyrfl
251	PNSKLSYAER	ELVLNANNYN	DMHINEKTES	KPKKSFRKPI	GKSKKHNLQI
			fs	™ =	
301	DFNSIDLSES	VIPGÇGFIPD	FSIHHLCKVP	ичччтында	LPLSFNTKNL
			X =		
351	NATSNESYLF	NDIVKIKSKS	IQKLVFNSDT	DNYHHTKYFY	TKTYRGPGSG
401	NYKDGALMNK	INKIELSSNK	KPFHKREVSN	nnrynkslkg	LVHEKFDKNP
451	VEYLLSEQRK	YTEDYSNLEI	LHNSLOFNVL	LUTYRGVAQE	TWNNYYKFKL
E01	TOPPOLYNIA	MESNO, DOOR	LDAARHQQWA	= X	fs = PLUPROPENT
201	The Bonestoo	MARKEL SARK			* X XX
551	FEQLQSEFGQ	RKKDLEEKLR	X X	£ 3	# = #3
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601			npapppgpie		
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551	missing	ALTOVIONE	IRMINIT	.FBRFRABIU	
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51	DDLAKVFRE	= OS TKKYPIKTY	ξ.			I	. a 6-
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1	COSYVPOSO	P XYSQQTQDR	G MFSGGGGGH	G HYQQQQQGYN	A YGPPPPQGG:	ť	
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51	YQQQPGGGG	AAUNUÜÜÜÜ	P MYVQQQPR30	GNESCTW&CI	AALCVCCTLE)	
	amb ==						
101	MLF				•	Fig	68
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22208							
		X X	X :=	R =	X		
1	MFDNFILKNL	VDETESEIDS	GETELSDDYY	YYYSYEDDGK	EDDSDEITAQ		
51	ILLSNSBLGT	XTPNFEOPFE	QINIEDNKVI	SVNTPKTAKP	TTTVFGTSTS		
	x		xx				
	=		82				
101	ALSTFESTIF	EIPKFFYGSR	rkqlssfknk	NSTIKFOVFD	WIFESGTTNE		
151	KVHGLVLVSS	GVLLGTCLLF	IL			Fig	69
						ry	0 /

		=			= =		
1	MRRREIERRA	: DOKKREGRÇK	EHEAKRDIRI	QQLSEQDSRS	NOTEXEEX	VF	
51	KKARSTNSGA	DETGLMSDKE	FDDSAYEPDY	LPZENLWNKP	NHPDTNHK	TK	
101	KYTENVVENL	DEPPNOTSAY	nssfhdetni	ONEIGIPEND	EYVPQMKA	TS	
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151	SVNNTTIPAQ	RRHESLSTSE	NKRRXFETAD			KI	
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201	QVSONPWATVY	ffmxxkrlet	PEGKLLCRDQ			Fi	g 70
328c1							
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1	MPRIKÇVDVF	TYTKYLGNPV	AVIYDSDMLT	WEALXQMEQT	TNLSETTFI	I.L.	
51	TPKSSIADYS	IRIFTSGGNE	LPFAGHPTLG	TAFALLEDGK	IK SNDNGO:	I	
		ambigui	ties				
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101	QECGAGLVKI	Svektpninn	nninnksnel	PFLLSFELPY	FKFHBIDDE	TV .	
		fs			G N		
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151	IBELQHSWNG	TNIIGKPVLI	Dagpkwavfq	rgsgkzvldi	NVDLAQIER	IL.	
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201	SLENGWIGIG	VFGKHNENGD	SVELRNIAPA	VGV		tig	<i>i </i>

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33qK part1

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1	NVTDST?F!	M GFLGSTFYA		===== Y DLATLHLLFV	SQTQTPSRIT	
	C XF	х				
51	יישטעענינג די א	e V nevenbrers	- DDCD* Fires			
J.	Amelian A	1 VERGENIETI	r rakokuman	V CEGNSTVNQL	LVFGEYLIAT	
		x	X X	-	x xxxx	
101	TLEGDIFVF.	≥ R KTEGKKFPTa	e e ELYTPIAITN	= LVEGEIVGLI	= ===== EDDTV1 NEX71	
			· ·	2.00111021	HEETITANA?	
	XXX X					
151	VATTQSVFV	e myrtgkllyk	SRELQFEGER	ISSIEAAPVL	DVIAVGTSNG	
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		=				
	3	-				
201	NVFLFNIKKG	_	LELMLPSKVA	SISFRTDGAP :	HLVAGLNINGD	
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251	LYFYDLDXKS	RVEVLENARK	ethggvanak	FLNGQPIVLS 1	nggdnhlkep	
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301	UPODNI PRKA	זעפפספונידפ	e perceues ne	= = VAIEFPQEDK 1	=======================================	
301	A. DENII. 1214	GETALETUNE	RINGGRIAFF	AWIELEAFDY J	THE LISASRU	
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	-	22			*******	
351	KTFWTFSLRK	Daqaqemsqr	LQK3KDGKRQ	agqvvsmrek f	PEIISISSS	
	dms					
401	YAR					
774	• m/				Fig	72

33qK part2

		fsG X	Xfs	XG FFXX	KE V S	DPTX		
	:	MLQNNRADLD	XVQNDIWYDVY	RKLKLDILNE	SSSSISEQIH	IRDRISRVYÇ		
		x	x		x x	X		
480c								
							<i>''</i> J	
	101	NETSLHEALE	ERK STNDEKK!	NKMDSLVKYC	ASIVSFIS		Fig	74
	51	SLNSFVPLTE	MTNFIQALNA	GLKENANYEI	WETLYAMFEN	ihgdvihofe		
	1	KLRKLDTNGN	Kafeseftkl	LPEAGESGQF	EFFLTYLLNL	SPAVLDLEIR		
33aK	part3					· · · · · · · · · · · · · · · · · · ·		
	351	NTLLHLDTIK	QQSKPKEAPK	KPENAPFFLQ	LTG		Fig	73 —
	251		LDGFLDEDSN	:		TLSSEPRSKF	_	
	201		ILATTHVSGN					
	151	ILYGHTMRIS	gmdfspdgrw	IVSVALDSTL	RTWELFTGGC	15GVILPIVA		
	101	dfgksvylgk	LQLEAPITSM	IYHKLSDLVA	CALDDLSIVV	IDVTTQKVIR		
	51	SLGGIGSYNL	QSCLLRKKYV	THKÖYALGTY	. iogáirknys	CGLDGIVGFY		
	i	IITAHKDETF	ARTWOSANKR	VGRHLLNTI	GGIVKSVCVS	QCGNFGLVGS		

*XFGXXXXIXQ*X X IXV missing sequence 101 KETPETIPLM MKELLQHQVQ IHQMQDKELD QLRVLIARQK QIGELINAEV missing sequence 151 EEQNEMLDRF NEEVDYTSSK IKQARRRAKK IL

51 PRILDLYRAI GTOMEEALKK KQLYSQLQES IDNLLVQEVP RSKRYLGGAV

		NS	elM X	XXGXN	R XFXW X	
				22 2 2 2		
1	NLTLKYIHF	/ KTREOTENAT	SHKSIETLEK	INYANVECMM	EDDLSSTSEI	
		GX X		Х	XX X	
	32 6 2	7= =		=	72 2	
51	IRRFTLEGVK	SQTSTSKDIT	SQÇKLDNFNT	ILRETREDEX	VVEDYLIDVI	
	F * X			R		
	= = =					
101		1:2001817		E		
101	APQIQLQSED	7552447731	PSIKGKILSI	RDSKNMANQI	LLETRYGILL	
151	KDANVFVLNK	FORESCHOOL	STEMPVENZE	Nicobult const	EUNIORN CAN	
	1012//11/21//	22170CIDAL	DIDNETONG	"WEENDG! DI	TONGKANGAN	
				A		
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201	nlliexLsvm	TMCYESEILS	SKLSFNAGDL	200	DNSKOAPLRL	
201	nlliexLsvm	TMCYESEILS		200	DNSKQAPLRL	
201	NLLIEKLSVM	TMCYESEILS	SKLSFNAÇDI FV	200	DNSKQAPLRL IB	
			FV ==	DQEEQENYND	IB	
	nlliex l svm Gidmpsvvit		FV ==	DQEEQENYND	IB	
	GIDMPSVVIT		FV ==	DQEEQENYND	IB	
			FV ==	DQESQENYND EPMSKVIHKK	IB	
251	GIDMPSVVIT * F	STSSQYFTLY	FV == VIIVSLLFYS E YXX	DQEEQENYND EPMSKVIHKK X * X = = =	IB 22 IEFMRPSIDF	
251	GIDMPSVVIT	STSSQYFTLY	FV == VIIVSLLFYS E YXX	DQEEQENYND EPMSKVIHKK X * X = = =	IB 22 IEFMRPSIDF	
251 301	GIDMPSVVIT F EBLGALTSRL	STSSQYFTLY	FV == VIIVSLLFYS E YXX	DQEEQENYND EPMSKVIHKK X * X = = =	IB 22 IEFMRPSIDF	
251 301	GIDMPSVVIT * F	STSSQYFTLY	FV == VIIVSLLFYS E YXX	DQEEQENYND EPMSKVIHKK X * X = = =	IB 22 IEFMRPSIDF	Fig 76

				FIG				
		*******		===	DI DECCE			
	1	ITEFSUFKIT	r Klpalaeldi	LKRCYICEDI	. INAPVETOCO	HTYCSQCIRE	i	
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5	1	FILRDNRCPL	. CKTEVFESGL	KRDFLLEETV	ISYASLRPHL	LRLLEIEKVE		
10	1	SKQEVDREKS	ANESALNGNR	N'AINDVDET	RVKDQLNADK	lgeekgqaqh		
		G fs				ж		
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15	1	Wegnegtte	VILLISDOBE	NGSDSLVKC?	ICFERMELDV	LQGKHIDDCL		
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20	1	SGKSTKRTPT	CILGPKAKAP	KÇITSFFKPT	IDTKTPSPPT	Skasttptat		
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25	-	PTITLLXAMV	aspspyaqst	AHKGKEFEKF	DFSSLSTQKI	KAKLSDLKLP		
			•	7				
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30:	1	TTGSRNEMEA	RYLHYYVIYN	anldsnhpv			Fig	' /
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;	1	MOFSSAVVLS	AVAGSALAAY	SNSTVTDIQT	T.VTITSCEZ	NKCHETEVTT		
51	1	GVTT/TEVT/T	TYTTYCPLST	TEAPAPSTAT	DVSTTVVTIT	SCEEDKCHET		
101	1 .	avtogvttvt	EGTTIYTTYC	PLPSTEAPGP	apstablishp	Aesepuptta		
151	1.	AESSPAKTTA	AESSPAQETT	PKTVAAESSS	aettapavst	aergaanav		
201	1	PVAAGLLALA	alf				Fig	78

x x

- 1 REESIISEEM KEGIPFFTIV ARIPVIEAFG FSEDIRKKTS GAASPQLVFD
- 51 GYDMLDIOPF NVPHTSEBLE ELGEFAEREN VARRYMUNIR RRKGLFVDEK

x x

101 VVKNAEKORT IKRD

Fig 79

29q3 part2

1	WRICQGSC	e pevhehlin	L IDSPGHIDF	S SEVSTSSRL	C DGAVVLVDVV	
51	EGVCSQTVN	V LRQCWIDKL	K PLLVINKID	R LITEWKLSP	L EAYOHISRII	
151	equnsvigs	f fagdrledd	L MWREAGSVG	E PIEKSDEDL	Y FTPEKNOWIF	
151	ASAIDGWAF	s voctaktyli	KLGFSQQAL:	s Ktlwgdfyl:	D MKNKKIIPGK	
201	KLKNNSNSL:	K PIFVSLILDO	VWAVYZNOVI	ERNQDKLEK	: ieklgakitp	
					XXX	
					202	
251	KDLRSKDYKI	n linlimsqwi	PLSHAILGSV	/ TEYLPSPIV	QRERIDHILD	
	XXX XXXX X	XX XXXX	Х	ambiguitie	:8	
	333 888 2 5			======================================		
301	ETIYSAVDSE	: LEKSKLVDPS	FVKAMQECDS	SHPETHTIAY	VSKLLSIPNE	
	fs			ambi	guities	
	=				255	
351	DLPKASNAAT	GGLTADEIQE	RGRIARELAX	Kaseaaalaq	EGSKNEDEFA	
	x					
.: 61	エリアリンスのサルム					
401	TERESTREM	efeeddfene	EDESDANAVE	ESTETIVGFT	RIYSGSL5RG	
		X	s	Y G	NFR	
		-	•		# 77 #	
451	QKLTVIGPKY	DRSLPROHOT	NFEQITNEVE	IKDLFLIMGR	BLVRMEKVPA	
		3	*R	ν		
		=	# LE	-		
501	GNIVGVVGLO	NAVLKNATIC	SPLPEDKPYI	NLASTSTLIH	NKPIMKIAVE	
551	PTNPIKLAKL	ERGL			<u></u>	CV
					Fig	80

1 PPKVAKSKES TIGKIFRYTF YTAVISVIGS AGLIGYRIYE ESQPVDQVKQ

X

51 TPLFPNGEKK KTLVILGEGW GAISLLKNLD TTLYNVVIVS PRNYFLFTPL

s X fa fa

101 LPSVPTGTVE LRSIIEPVRS VTERCPGQVI YLEASATNIN PKTNELTLKQ

R N XX

151 STTWYSGHSG RETSSSKSTV AEYTGVEZIT TFLNYDYLVV GVGAQTILIF

201 GNPGERMRKF NPFFERRUSG SHLQIR

Fig 81

357cL

1 MAKFIKAGKV ALVVRGEYAG KKVVIVKFHD EGTKSHPFPH ALVAGIERAP

51 LKVTKKMDAK KVTKRTKVKP FVKLVNYNHL MPTRYSLDVE SFKSAVTSEA

is

101 LEEPSQREEA KKVVKKAFEE KHQAGKIMWF FQKLHF

1 MSIPSTQYGF FYNYASGLNI KKOLPDNKPG AGQLLLKVDA VGLCHSDLHV X fs D X X X 51 LYEGLDCGDN YVMGHELAGT VAELGEEVSE FAVGDRVACV GPMGCGLCKH missing sequence ======== ambiguitles X XXXXIFGXXEXX -----101 CLTGNONVCT KSFLOWFOLG YNGGYEQFLL VKRPRNLVKI PDNVTSERAA missing sequence 151 AITDAYLTTY HAIKSAGVGP ASNILIIGAG GLGGNAIQVA KAFGAKVTVL missing sequence 201 DKKDKARDQA KAFGADQVYS ELPDSVLPGS FSACFDFVSV QATYDLCQKY missing sequence 251 CEPKGTIVFV SLGATSLNIN LADLDLREIT VKGSFWGTLM DLREAFELAA missing sequence 301 QGKYKPNYAH AFLSELPKYM EKLRAGGYEG RVVFNP

Fig 83

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1	MEKIDINTN	S NKIQQAYDKV	VRGDENATFV	VYSVDKNATM	DVTSTGDGSL	
	¥	P			P LSYD	
51	= EDFVEHFTD	= G @VQFGLARVI	VPGSDVSKNI	LLGWCPDSAP	arlrcafann	
	, .	**	N W "	y		
	3 I	M =	= = = N A H	Х =		
101	FADVSRVLS	G YHVQITARDQ	DOLLYNEFLN	RVGAAAGARY	Stotsglkk?	
151	SPAAPKPTS:	k fyvaksesas	KPSFVPKSTG	KPVAPAKPKP	Knitkdagme	
201	DAEDVEERD:	P DKKPLDNV3S	AAKNIKAWID	ELPKOKSDET	SSTPKTFKSE	
•		?	HD R	T		
251	PQEEKNDDD	G ÇSKPLSERMK	AYDQPSSSDG	RLTSLPK?KI	GHSVADKYKA	
		Ís				
301	er eanar r bi	= A FGAKPAFGTO	יוזארצשטרנוט	GGLSRDFGAE	NGKTPACTWA	
201		_				
351	EKRGXYKTV	A EDEKETNSSE	KVDEPSEHHA	adlakkfeek	Aniagdipsl	
			ĸ	s fs	х	
401	PTRNLPPAPI	ARETAIPSNE	KOKEEKSEEE	Qapapslptr	NLPP?SQRQ?	
	fs X	ХS		missim	sequence	
453	= =	uu Eesebeapap				
451	epa teyee e	i EELEDEATAF	SHENDANDELV	Parenessan	Antiumin	
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501		FSEGDLIIDI				F19 84
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226c af2			······································		~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~~	
1	VLGSQWGDEG	KGKTADTTCD	DODRACVCIC	GNNAGHTIVV	GKVKYDFHML	
51	PSGLVNPKCQ	NLVGSGVVIH	VPSFFAELEN	LEAKGLDCRD	RLFVSSRAHL	•
	A	D IX				
101	AND SEVENDE	= == LKEAELSTNK	KSTGTTGKGT	CPTYSTRASE	SGIRVHHLVDI	•
	-					
151	POPEAWEEFK	TRYLEL/ESR	QKRYGEFEYD			I. 25

1 EDKKORFDAS GASAVODKTA TAILRRKKKI NALVVDDATN DONSVITMAS	EDKKQSFDA	AS GASAVODKTA	TAILREKKET	MALVUDDATM	DOMENTOME
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- 51 NIMELLQLER BDIVLVWGKK RESTVLIVLA DDDMPDGVAR VARCVRNNLR
- 101 VRLGDIVTVE FCPDIKYANR ISVLFIADTV EGINGSIFDL YLKPYFVEAY
- 151 RPVFKGDLFT URGGMRQUEF KUVEYDPEZI AIVAQDTIIH CEGEPINRED

F fs

- 201 SENSINEVGY DDIGGCKKOM AQIRELVELP LRHPQLFKSI GIKPPKGILM
- 251 YGPFGTEKTI MARAVANETG AFFFLINGPE IMSKMAGESE SMLRKAFEZA

fs ? fs NKK P

301 EKNSPSLIFI DEIDSIAPKR DKTNGSVERF VVSQLLTIMD GMKARSNAVV

G DQ L Pfs

- 351 IAATMRPNSI DPALRRFGRF DREVDIGVPD AEGRLEILRI HTKNMKLADD
- 401 VDLEATASET HGFYGADIAS LCSEAAMQQI RERMDLIDLE EETIDTEVLN
- 451 SLGVTQDNFR FALGNSNPSA LRETVVENVN VTWDDIGGLD MIKNELKETV
- 501 EYPVLHPDQT CKFGLAPTKG VLFFGPPGTG KTLLAKAVAT EVSANFISVK
- 551 GFELLSMWYG ESESNIRDIF DKARAAAPTV VFLDELDSIA KARGGSHGDA
- 601 GGASDAVVNQ LLTENDGMNA KKNVFVIGAT NRPDQIDPAL LRPGRLDQLI
- 651 YVPLPDEPAR LEILQAQLEN TPLEPGLDLM: EIAKITHGFS GADLSYIVQR

AfsQ I

- 701 SAKFAIKOSI EAQVKINKIK EEKEKVKTED VOMKVOEVEE EDPVPYITRA
- 751 HFEEAMKTAK RSVSDAELRR YESYAQQLQA SRGQFSSFRF NENAGATONG

X

801 SAAGANSGAA FON

Fig 86

£s.

1	TLKCRLEEI:	L fakagevkof	KKEHGKTVIO	EVLLEDAYGO	MRGIKGLVWE		
51	GSVLDPIEG	I RFRGRTIPDI	QKELPKAPGG	EEPLPEALFW	LLLTGEVPTD		
101	AQTKALSEZI	F AARSALPKEV	EELIDRSPSH	LHPMAQFSIA	VTALESESOF		
151	aqayakgani	(SEYWKYTYED	SIDLLAKLPT	IAAKIYRNVF	HOGKLFAAID		
201	skldyganla	X T = = SLLGFGDNKE	=	IHSDHEGGNV	T SAHTTHLVGS		
251	Alsspeisla	ycingrygji	RI * == = HGRANQEVL			Fig	87
9903 part2							
1	QREFALKHMP	DYELFKLVS27	iyevapgylt	KHGKTKNPWP	nvdshsgvll		
		S =					
51	QYYGLTEQSF	YTVLFGVFRA	FGVLPQLILD	rgigmpier?	KSFSTÉKYIE		
101	LVKNINK					Fig	88
409c5 part							
	ambiguityX		X	, GT			
1	SDYHVIWLAR	RNNGIMEAEY	RLYLLVITLI	ispvsximfg	vgaarewpwq		
51	VIYVGLGFIG	FG%GSIGDTS	Cqyadmiyem	IVIQGMVGVS	IINNTLACIF		
101	TFACSYVILDG	SCTQNTYIAL	SIIDFATTAL	V FPPLYYGKT	FRRKTKRLYV		
151	SMVELTQGM					Fig	89

ix

409c5 part2

1	DQNNEDFIPG	TUNITSLEVD	SEDENVSHYD	ASSRPKVKTK	GNYTTEDADO
-	28 25. 21.3			WOODLEVALTY	

- 51 NSCNDPLNWS KWRKLSNPFI VIFITAFTAA TSNDAGSIQD SLNEKYGISY
- 101 DAMNTGAGVL FLGIGWGTFF LTPASSLYGR KITYFICIFL GLLGAVWFAL
- 151 VKSISDSIWS QLFVGISESC AEAQVQLSLS ELYFAHNLGS VLTSYIVATS
- 201 VGTYLGPLIA AFIVONIGFR WVGWIAAIIS GALLFVIVFC LDETYFDRAK
- 251 FTKP

F19 90

4 to 2

ESP1

- 1 DEGLEDILHE VESKWEGGEI SGIFTNDNDV ENESKNYTHK EKODLMKILK
- 51 DOLTVSDDKS NIERFLQFNI FIYYOFYSME EYNYELVDDL IXFITINMNS

Х

101 HGRIVNFGIN VEINKLHELI KNIIDKVNKN KCRCDXQQQK QQQQQQQQQ

esitiugidme

22QQ fs X ambiguities

- 151 NSNNSQHIVL IFNANCSNFP WESMEFLREK SISRMPSIHM LLDLVKSMTN
- 201 NKNKLMFVDK SNLYYLINPS GDLIRSENRF KKLFESNHLW RGEIGKLSSN

F

X ambiguities

- 251 EHEDYQDSIL CEILKSHLFV YIGHGGCDQY IKVSKLFKKC GMNQDLLNKL
- 301 PPSLLLGCSS VKLENCHYNY NSSMLQPLGN IYNWLNCKSS MILGNLWDVT

D

- 351 DKDIDIFTLS LUDKWGLIAU YNGSGHDYGM KKLDLTNOVV QSRSKOTLKY
- 401 LNGSAPVVYG LPX

-19 91

The see in

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1	STRGIKVHAT	FESKNERPUL	TYGIIAIGE	TARMIÑE KWT	WOT IRENDAT
					M
51	AQAQSGTGKT	ATFSIGNLEV	idikskecqa	LILSPTRELA	iðiðmaakht
161	GDYMVISTHA	CIGGRAVGED	AKKT ÖĞGĞĞİ	VSGTPGRVID	VIKRRNLOTA
151	HIKVLILDEA	Delftngfke	OIAEIAKHT Ł	PSVQVVVVSA	TLPREVLENT
	fs		T .		
201	= SKFTTD9VKI	LVKRDEISLL	GI KĴYYV Ĵ CE	REDWKFDTLC	DLYDNLTITQ
251	AVIFONTALK	VNWLADQMKK	QNFTVVAMHG	DMKQDERDSI	MIDFFRGNSR
301	VIISTEVWAR	GIDVÇQVSLV	INYDLƏTOKE	NYIHRIGRSG	rfgrk gt ain
		x			
351	LITEODVVTL	YELEKYYSTK	IKEMPMNIND	IM	Fig

40c at

1 NVDTOLITET RFILQEQQTV AFTATGELSL LLNALQFAFK FIAHWIRRAE

51 LVNLIGVSGS ANSTEDVQXX LUVIGDEIFI NAMRSSNNVK VLVSEEQEDL

101 IVFPGGGTYA VCTTPIDGSS NIDAGVSVGT IFGVYKLQEG STGGISDVLR

missing requence

75224222224282E43

151 PGKEMVAAGY TMYGASAHLA LTTGHGVNLF TLDTQLGEFI LTHFNLKLPD

missing sequence

201 TKNIYSINEG YSNKFPEYVQ DYLKDIKKEG YSLRYIGIMV ADVHRTLLYG

missing sequence

251 GIFAYFTUKL RVLYEOFPMA LLMEQAGGSA VTIKGERILD ILPKGIHDKS

missing sequence ***************

301 SIVLGSKGEV EKYLKHVPK

Fig 93

92

1 DNVSSTSTAE AVMNEIKVKD EFPQEEQAHT SLEDKPV	SAY IGIIIMOFLI
51 AFGGFVFGFD TGTISGFINN SDFLERFGGT KADGTLY	FSN VRTGLMIGLF
x x	
101 Naccatoria isusanuon nuotimatti taisa	
101 NAGCAIGALF ISKVEDMYGR RVGIMTAMIV YIVGIIVQ	ia sqhawyqvmi
	ambiguities
	≠= X
151 GRIITGLAVG MLSVLCPLFI SEVSPKHLRG TLVCCPQL	T
TOT GRITTOMAN MASVACETET SEVSPREERG TEVECTOR	MI TEGIFLGYCT
fs	
201 TYGIKSYSDS RQWRIPLGLC FAWALCLVAG MVRMPESP	RY LVGXDRIEDA
	2R ==
251 KMSLAKTNKV SPEDPALYRE LQLIQAGVER ERLAGKAS	G TLFNGKTKIF
IV missing sequence	
## ###################################	Fig. 94
301 BRVMLGVMLQ ALQQFNWGKN LFPSYLTSXP N	rig it
98c cp	
	ng sequence
1 NAPVSGTITE FLYDYDATVE VGQETIKMEE GDAPAGGAS	
· minoi	
	ng sequence
51 PEKAKEBSAP AAAPKKESTK KEEPKKESKP APKKEESKK missing sequence	S TOSTOSAPTF
51 PEKAKEESAP AAAPKKESTK KEEPKKESKP APKKSESKK	S TOSTOSAPTF
51 PEKAKEBSAP AAAPKKESTK KEEPKKESKP APKKEESKK missing sequence	S TOSTOSAPTF
51 PEKAKEBSAP AAAPKKESTK KEEPKKESRP APKKEESKK missing sequence 101 TNFSPNEERV KMIRMRLRIA ERLKESQNTA ASLTTFNEVI missing sequence	S TOSTTSAPTF D MSNLMDFRKK
51 PEKAKEESAP AAAPKKEETK KEEPKKESKP APKKEESKK missing sequence 101 TNFSPNEERV KMIRMRURIA ERUKESQNTA ASLTTFNEVI missing sequence 151 YKDEFIEKTG IKLGFMGAFS KASALALKEI FAVNAAIEW	S TOSTTSAPTF D MSNLMDFRKK
51 PEKAKEESAP AAAPKKEETK KEEPKKESKP APKKEESKX missing sequence 101 TNFSPNEERV KMIRMRLRIA ERLKESQNTA ASLTTFNEVI missing sequence 151 YKDEFIEKTG IKLSFMGAFS KASALALMEI FAVNAAIEM missing sequence X XX XX XX	S TOSTTSAPTF D MSNLMDFRKK N DTLVFKDYAD
51 PEKAKEESAP AAAPKKEETK KEEPKKESKP APKKEESKK missing sequence 101 TNFSPNZERV KMIRMRLRIA ERLKESQNTA ASLTTFNEV missing sequence 151 YKDEFIEKTG IKLGFMGAFS KASALALKEI PAVNAAIEM missing sequence X XX XX X	S TOSTTSAPTF D MSNLMDFRKK N DTLVFKDYAD
51 PEKAKEESAP AAAPKKEETK KEEPKKESKP APKKEESKX missing sequence 101 TNFSPNEERV KMIRMRLRIA ERLKESQNTA ASLTTFNEVI missing sequence 151 YKDEFIEKTG IKLGFMGAFS KASALALMEI PAVNAAIEM missing sequence X XX XX XX 201 ISIAVATPKG LYTPVVRNAE SLSILGIEKE ISNLGKKARI	S TOSTTSAPTF D MSNLMDFRKK N DTLVFKDYAD
51 PEKAKEBSAP AAAPKKESTK KEEPKKESKA APKKEESKA missing sequence missing sequence missing sequence 151 YKDEFIEKTG IKLGFMGAFS KASALALKEI PAVNAAIEM missing sequence 251 XXX X C X X X F XF XF X IX	S TQSTTSAPTF D MSNLMDFRKK N DTLVFKDYAD NX * C GKLTLEDMTG
51 PEKAKEBSAP AAAPKKEETK KEEPKKESKP APKKEESKX missing sequence 101 TNFSPNEERV KMIRMRLRIA ERLKESQNTA ASLTTFNEVI missing sequence 151 YKDEFIEKTG IKLGFMGAFS KASALALMEI PAVNAAIEM missing sequence X XX XX X 201 ISIAVATPKG LVTPVVRNAE SLSILGIEKE ISNLGKKARI S X XX X C X X X F XF X X X IX	S TQSTTSAPTF D MSNLMDFRKK D DTLVFKDYAD NX * C GKLTLEDMTG

249c at

- 1 ERMISGMIYN CLQKELETTR MSCRDYMLLY GSFRTRDYKT TQEFLDAKYK
- 51 HLESPIGHVG KNAFMEYPIY FDYGFNTYLG DNFYSNYNLT ILDVSIVRIG

fs

101 NEVKOGPNYS ILIPTHPVDP TLRYDQLENA LPVTVGNGVW LOGSCTILGG

X missing sequence

151 VTVGDGSIVA AGAVVNKDVP PNTVVAGVPA RVVKQLEPRD PNF

Fig 96

55a1

XX XXFY X P X X

1 TSDTKTKQRB NELLKDISSQ GGMLRTVPRS SSSSSQKKK SSKKQRHNDE

T HL V L

51 DDZENGGGEG FLDASSSRKI LQLAKEQQDE LEQEDEIQNK PSFAQSFKNQ

C L FI S

101 QIDSEEEEE DEYSDFERER EVEETVYDEE DAEVDPKDAE LENKYFQSNG

151 EANENDEDNS FORTNIADK ILAKIQEKES QQQQQQQSSP DNSNEDAVLL

E I

201 PPKVILAYKK IGQILSTYTH GKLPKLFKIL PSLKNMQDVL YVTNPNSWTF

251 HATYEATKLE VSNLSSNEAT VFTETILLPR FROSIENSOD ESLNYHIYRA

301 LKKSLYKPGA FFKGFLLFLY DGYCSVREAT IAASVLTKVS VPVLHSCHYC

351 GVLMNKRES EVEVLREI

•	1 MIVYKADORK EPVREDKITA RVQRLCYGLN PNHVEPVAIT QKVISGVYQ
3;	VYTTELDNIA AEIAATMITTI HPDYAVLAAR IAVSNIHKQT TKQYSKVSK
101	L LYEYINPKTG LHSPMISKET YDIIMEHZDE LNSAIVYDRD FNYNYFGFK
153	LERSYLLRIN GRVAERPOHL IMRVAVGING NDIPRVIETY NLMSQRFFT:
301	GSPCLFNAGT FRFQMSSCFL LAMKDDSIEG IYDTLKSCAL ISKSAGGIGI
251	HIHNIRSTGA YIAGTNGTSN GIIPMVRVFN NTARYVDQGG NKRPGAFALY
301	LEPWHSDIFD FIDIRKNHGK BEIRÄRDLFP ALWIPDLFMK RVEQNGDWTL
351	FSPNEAPCLA DVYGDEFEEL YTKYEKENRG RQTIKAQKLW YAILGAQTET
401	GTFFMLYXDS CHIKSNOKNL GIIKSSNLCC EIVEYSAPDE VAVCNLASIA
451	LPSFVENDSK STWYNFORLH QVTKVVTRNL HRVIDRNHYP VPEAERSKMR
501	HRPIALGVQG LAJAFMELRL PFDSQEAREL NIQIFETIYH AAVEASIELA
551	KEEGAYETYP GSPASQGLLQ FDLWNRKFTE LWDWDTLXQD LAKHGMRNSL
601	LVAPMPTAST SZILGNNECF EPYTSNIYSK RVLAGEFQIV NPYLLKDLVD
651	LGVWNDAMKS SITANNGSIQ ALPNIPDEIK ALZKTVWEIS QKHIIDMAAD
701	RAAFIDQSQS LNIHIKDPTN GKLTSMHFYG WKKGLKTGMY YLRTQAASAA
751	IQFTIDQKIA ETAGHIVANL DKLNIKKYVN KGRVESENTS DAPYKSFSTE
861	PTSLESSVAD LKEKDEGEKF AEDKTIEELE NDIYSAKVIA CAIDNPESCT
851	MCSG

F19 98

6 F. J

485cL

fs X fs V

- 1 APKWYQSEDV PAFKQTRKTA RPQKLRASLA PGTVLILLAG RFRGKRVVYL
- 51 KNLEDNTLLV SGFFRVNGVP LRPVNARYVI ATSTKVNVSG VDVSKFNVEY
- 101 FAREKSSKSK KSEAEFFNES OPKKEIKAER VADOKSYDAA LLSEIKKTPL
- 151 LKQYLAASFS LKNGDREHLL KF

ال القطية ال

1 MENDKGQLVE LYVPRKCSAT MRIIKAKDHA SVQISIAKVD EDGRAIAGEN Fig 100 51 ITYALSGYVR GRGEADDSLN RLAQQDGLLK NVWSYSR 67a1 part1 X 1 MSFKGFKKGV LRAPQTMRQK FNMGEITQDA VYLDAERRFK EIEMETKKLS 51 EESKKYFNAV NGMLDEQIDF ARAVAEIYKP ISGRLSDPSA TVPEDNPQGI 101 EASELYQAVV KOLKOTLKPO LELIENRIVE LAQELLKIIQ AIRKMS 67q1 part2 1 HKRNFSKYEL KKERTVKDEE KMFSAQTEVE IAQQEYDYYN DLLKNELPVL Z. Y D 51 FOMOSOFIKP LEVSEYYMOL NIFYTLYTEM BELKIPYFOL STDIVEAYTA s fs K 101 KKGNIEEQTO AIGITHFKVG HAKSKLEATK RRHAAMNSPP PTGASSIAST 151 GTGGELPAYS PGGYNQPYGD SKYQPPSSPA TYQSPVVAAT AQSPATYQSP 201 VATGOPPSYL POTPASAPEP OVGSGLPTCT ALYDYTAGAG GOLTFPAGAV

251 IEIIQRTEDA NGWYTGKYNG QTGVFPGNYV QL

1	Mitsketflf	TSESVGZGHF	DKICDQVSDA	ILDACLAVDP	LSKVACETAA

- 51 KTGMIMVFGE ITTKAQLDYQ KIIRDTIKHI GYDDSEKGFD YKTCNVLVAI
- 101 EQQSPDIAQG LHYEKALBEL GAGDQGIMFG YATDETDEKL PLTILLAHKL
- 151 NAALASARRS GSLFWLRPDT KTQVTIEYEK DGGAVIPKRV DTIVISTQHA
- 201 EEIPTEMIRK EIIEHIIKQV IPEHLLODKI IYHIQPSGRF VIGGPQGDAG
- 251 LTGRKIIVDT YGGMGAHGGG AFSGKDFSKV DRSAAYAARN VAKSLVTAGL
- 301 AKRALVÇESY AIGVAEPTSI YIDTYGTSKL STEALVEIIK NNFDLRPGVI
- 351 VKELDLARPI YFKTASYGHF TNQENSWEQP KKLKF

Fig 103

.. 5" b U"

232c cp

- 1 PKNSADAKYT METIRGIRLY GKALKIKRID AKSQSSTNNP NNQTIGTFVQ
- 51 SDLINPNYID VGAKLFINNL NPLYDESFLM DTFSKFCTLI RNPIIRRDSE

T missing sequence

101 GHSLGYGFLT YDDFESSDLC IQEARNTILM NNKIAISYAF KDLSVOGRKS

missing sequenc

151 RHGDQVERKE GXXXVP

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1	CPFLTE	eislh	TICHOHORINIY	INTAČHIL N	ED FA	IIILMD	нь Есс	DLFTI	III		
	\$ *) =								
51			RKVPRTOFET		LO LIE	CAIEYC	HE NNI	YHCDI	КÞ		
				•	•						
101	ENIMUR	MIPY	YVESTINNNN	NNGEDDLC	ya nsi	ICYNE	LH LVI	IDFGL	AM		
151	DSATIC	CNSC	RGSSFYMAPE	RTTNYNTH	RL INC	LIEMNO	OY ESI	EINGT	my.		
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251	72/MNKY.	CLSK	IIPISSQFNR	ILDRIFKL	ip ngr	IGL				• 5	
CD2											
, VIVA											
1	NGTKÇÇ	IET	idninkadlp	KDAEAAICE	P ALY	LGLAVE	O MKO	PTVAI	GA		
						x	•				
						•	,				
51	QNVFDKS	SCGA	FTGETCASQI	LDVGASWTI	T GHS	errtii	K ESD	EFIAE	KT		
			X	X	X =	GCWFQ		_	3X		
101	KF21.77G	UKV	= :lcigetlee								
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